TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 1 of ?)

Biomarker: Total Nicotine (ug/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	2388.230, 1566.130	5.121, 2.321	
	SD, SE	1770.0800, 250.3270	13.2003, 1.6248	
	MEDIAN	2032.230	1.980	
	MIN, MAX	22.137, 6337.875	0.530, 88.160	
	5%, 95%	152.361, 5830.270	0.800, 17.248	
	BLQ Values Included	0	44	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	2506.710, 1693.850	7.945, 2.295	
	SD, SE	1845.8400, 263.6920	42.6331, 5.2478	
	MEDIAN	2099.050	2.088	
	MIN, MAX	10.710, 9552.459	0.625, 348.555	
	5%, 95%	192.780, 5810.680	0.815, 7.128	
	BLQ Values Included	0	51	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	2737.440, 1824.200	2.766, 2.232	
	SD, SE	1983.6400, 292.4720	2.6681, 0.3284	
	MEDIAN	2327.150	2.082	
	MIN, MAX	10.061, 7734.320	0.750, 19.760	
	5%, 95%	288.769, 7314.620	0.990, 7.224	
	BLQ Values Included	0	52	
Overall*	N	50	66	0.5430
	MEAN, GEO MEAN	2497.610, 1731.920	5.278, 2.469	
	SD, SE	1659.2700, 234.6560	17.1813, 2.1149	
	MEDIAN	2046.090	2.147	
	MIN, MAX	14.303, 6909.320	0.967, 138.186	
	5%, 95%	172.571, 5294.820	1.057, 11.448	

[a] P-value from wilcoxon rank-sum test.
* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 2 of ?)

Biomarker: Total Nicotine (nmol/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	14723.500, 9655.180	31.572, 14.306	
	SD, SE	10912.6000, 1543.2700	81.3796, 10.0171	
	MEDIAN	12528.700	12.207	
	MIN, MAX	136.475, 39072.999	3.267, 543.506	
	5%, 95%	939.306, 35943.600	4.932, 106.334	
	BLQ Values Included	0	44	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	15453.900, 10442.600	48.983, 14.152	
	SD, SE	11379.6000, 1625.6600	262.8330, 32.3525	
	MEDIAN	12940.600	12.870	
	MIN, MAX	66.027, 58890.910	3.853, 2148.842	
	5%, 95%	1188.490, 35822.800	5.024, 43.944	
	BLQ Values Included	0	51	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	16876.300, 11246.200	17.055, 13.757	
	SD, SE	12229.2000, 1803.0900	16.4485, 2.0247	
	MEDIAN	14346.900	12.836	
	MIN, MAX	62.026, 47682.083	4.624, 121.820	
	5%, 95%	1780.260, 45094.600	6.103, 44.536	
	BLQ Values Included	0	52	
Overall[*]	N	50	66	0.5430
	MEAN, GEO MEAN	15397.800, 10677.300	32.537, 15.219	
	SD, SE	10229.4000, 1446.6500	105.9230, 13.0382	
	MEDIAN	12614.200	13.235	
	MIN, MAX	88.176, 42596.000	5.960, 851.919	
	5%, 95%	1063.900, 32642.600	6.514, 70.579	

[a] P-value from wilcoxon rank-sum test.

[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

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TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 3 of 7)

Biomarker: Total Nicotine (ug/mg Creatinin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value*
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	2.020, 1.298	0.003, NA	
	SD, SE	1.7387, 0.2459	0.0075, 0.0009	
	MEDIAN	1.727	0.002	
	MIN, MAX	0.021, 9.353	0.000, 0.044	
	5%, 95%	0.100, 5.254	0.000, 0.011	
	BLQ Values Included	0	44	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	1.974, 1.307	0.007, NA	
	SD, SE	1.6781, 0.2397	0.0408, 0.0050	
	MEDIAN	1.686	0.002	
	MIN, MAX	0.012, 7.495	0.000, 0.333	
	5%, 95%	0.144, 6.431	0.000, 0.004	
	BLQ Values Included	0	51	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	2.100, 1.463	0.002, NA	
	SD, SE	1.3912, 0.2051	0.0017, 0.0002	
	MEDIAN	2.056	0.002	
	MIN, MAX	0.009, 5.794	0.000, 0.011	
	5%, 95%	0.200, 4.966	0.000, 0.005	
	BLQ Values Included	0	52	
Overall*	N	50	66	0.1324
	MEAN, GEO MEAN	2.027, 1.398	0.004, NA	
	SD, SE	1.4655, 0.2073	0.0154, 0.0019	
	MEDIAN	1.811	0.002	
	MIN, MAX	0.014, 6.341	0.000, 0.126	
	5%, 95%	0.156, 5.620	0.000, 0.008	

* P-value from wilcoxon rank-sum test.
* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 4 of 7)

Biomarker: Total Nicotine (nmol/mg Creatin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	12.456, 8.001	0.021, NA	
	SD, SE	10.7192, 1.5159	0.0463, 0.0057	
	MEDIAN	10.644	0.012	
	MIN, MAX	0.129, 57.661	0.000, 0.271	
	5%, 95%	0.617, 32.391	0.000, 0.068	
	BLQ Values Included	0	44	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	12.170, 8.060	0.044, NA	
	SD, SE	10.3455, 1.4779	0.2515, 0.0310	
	MEDIAN	10.394	0.009	
	MIN, MAX	0.074, 46.207	0.000, 2.053	
	5%, 95%	0.888, 39.647	0.000, 0.025	
	BLQ Values Included	0	51	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	12.945, 9.019	0.012, NA	
	SD, SE	8.5770, 1.2646	0.0102, 0.0013	
	MEDIAN	12.673	0.012	
	MIN, MAX	0.055, 35.720	0.000, 0.068	
	5%, 95%	1.233, 30.615	0.000, 0.031	
	BLQ Values Included	0	52	
Overall[*]	N	50	66	0.1429
	MEAN, GEO MEAN	12.496, 8.617	0.026, 0.011	
	SD, SE	9.0349, 1.2777	0.0952, 0.0117	
	MEDIAN	11.165	0.010	
	MIN, MAX	0.086, 39.094	0.002, 0.779	
	5%, 95%	0.960, 34.649	0.002, 0.049	

[a] P-value from wilcoxon rank-sum test.
[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminating all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Free Nicotine (ug/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	1745.040, 1070.290	4.765, 2.123	
	SD, SE	1430.2000, 202.2600	13.2934, 1.6363	
	MEDIAN	1417.230	1.825	
	MIN, MAX	18.048, 5245.860	0.530, 93.100	
	5%, 95%	83.814, 4727.380	0.800, 13.992	
	BLQ Values Included	0	52	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	1806.110, 1170.200	6.987, 2.157	
	SD, SE	1358.1100, 194.0160	37.0042, 4.5549	
	MEDIAN	1573.750	2.013	
	MIN, MAX	10.980, 5904.527	0.625, 302.670	
	5%, 95%	94.792, 4729.200	0.815, 6.758	
	BLQ Values Included	0	55	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	2033.500, 1264.500	2.395, 2.049	
	SD, SE	1625.9700, 239.7360	1.8255, 0.2247	
	MEDIAN	1632.770	2.015	
	MIN, MAX	5.450, 6614.080	0.750, 13.650	
	5%, 95%	161.699, 5668.540	0.990, 4.740	
	BLQ Values Included	0	58	
Overall[*]	N	50	66	0.5246
	MEAN, GEO MEAN	1827.980, 1226.590	4.716, 2.293	
	SD, SE	1277.8800, 180.7190	15.1522, 1.8651	
	MEDIAN	1696.910	2.147	
	MIN, MAX	11.493, 4824.560	0.793, 121.310	
	5%, 95%	130.092, 4297.930	0.980, 8.342	

[a] P-value from wilcoxon rank-sum test.

[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2

STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 6 of ?)

Biomarker: Free Nicotine (nmol/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value*
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	10758.200, 6598.370	29.377, 13.087	
	SD, SE	8817.1700, 1246.9400	81.9539, 10.0878	
	MEDIAN	8737.190	11.251	
	MIN, MAX	111.266, 32340.727	3.267, 573.962	
	5%, 95%	516.713, 29144.300	4.932, 86.261	
	BLQ Values Included	0	52	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	11134.700, 7214.280	43.078, 13.297	
	SD, SE	8372.7600, 1196.1100	228.1310, 28.0810	
	MEDIAN	9702.170	12.407	
	MIN, MAX	67.692, 36401.409	3.853, 1865.961	
	5%, 95%	584.393, 29155.500	5.024, 41.663	
	BLQ Values Included	0	55	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	12536.500, 7795.640	14.764, 12.630	
	SD, SE	10024.1000, 1477.9700	11.2539, 1.3853	
	MEDIAN	10066.000	12.423	
	MIN, MAX	33.599, 40775.803	4.624, 84.152	
	5%, 95%	996.874, 34946.500	6.103, 29.222	
	BLQ Values Included	0	58	
Overall*	N	50	66	0.5246
	MEAN, GEO MEAN	11269.500, 7561.890	29.073, 14.137	
	SD, SE	7878.1100, 1114.1300	93.4135, 11.4984	
	MEDIAN	10461.500	13.235	
	MIN, MAX	70.852, 29743.400	4.891, 747.878	
	5%, 95%	802.015, 26496.800	6.042, 51.426	

* P-value from wilcoxon rank-sum test.

* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

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TABLE 14.1.1-2

STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Free Nicotine (ug/mg Creatinin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	1.551, 0.887	0.003, NA	
	SD, SE	1.6269, 0.2301	0.0074, 0.0009	
	MEDIAN	1.072	0.002	
	MIN, MAX	0.017, 8.892	0.000, 0.044	
	5%, 95%	0.071, 5.001	0.000, 0.012	
	BLQ Values Included	0	52	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	1.459, 0.903	0.006, NA	
	SD, SE	1.3786, 0.1969	0.0354, 0.0044	
	MEDIAN	1.262	0.001	
	MIN, MAX	0.012, 6.916	0.000, 0.289	
	5%, 95%	0.071, 4.633	0.000, 0.004	
	BLQ Values Included	0	55	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	1.565, 1.015	0.002, NA	
	SD, SE	1.1600, 0.1710	0.0013, 0.0002	
	MEDIAN	1.471	0.002	
	MIN, MAX	0.005, 5.060	0.000, 0.008	
	5%, 95%	0.107, 4.177	0.000, 0.004	
	BLQ Values Included	0	58	
Overall[b]	N	50	66	0.1132
	MEAN, GEO MEAN	1.522, 0.990	0.004, NA	
	SD, SE	1.2602, 0.1782	0.0136, 0.0017	
	MEDIAN	1.351	0.002	
	MIN, MAX	0.011, 5.866	0.000, 0.111	
	5%, 95%	0.119, 4.066	0.001, 0.006	

[a] P-value from wilcoxon rank-sum test.
[b] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 8 of ?)

Biomarker: Free Nicotine (nmol/mg Creatin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	9.559, 5.466	0.020, NA	
	SD, SE	10.0298, 1.4184	0.0455, 0.0056	
	MEDIAN	6.606	0.009	
	MIN, MAX	0.105, 54.819	0.000, 0.271	
	5%, 95%	0.438, 30.831	0.000, 0.074	
	BLQ Values Included	0	52	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	8.998, 5.566	0.039, NA	
	SD, SE	8.4990, 1.2141	0.2182, 0.0269	
	MEDIAN	7.780	0.006	
	MIN, MAX	0.074, 42.637	0.000, 1.782	
	5%, 95%	0.438, 28.562	0.000, 0.025	
	BLQ Values Included	0	55	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	9.650, 6.259	0.011, NA	
	SD, SE	7.1517, 1.0545	0.0080, 0.0010	
	MEDIAN	9.066	0.009	
	MIN, MAX	0.031, 31.195	0.000, 0.049	
	5%, 95%	0.660, 25.751	0.000, 0.025	
	BLQ Values Included	0	58	
Overall[*]	N	50	66	0.1274
	MEAN, GEO MEAN	9.382, 6.109	0.023, NA	
	SD, SE	7.7691, 1.0987	0.0834, 0.0103	
	MEDIAN	8.328	0.010	
	MIN, MAX	0.070, 36.164	0.000, 0.682	
	5%, 95%	0.734, 25.067	0.004, 0.039	

[a] P-value from wilcoxon rank-sum test.
[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 9 of ?)

Biomarker: Total Cotinine (ug/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	4878.380, 3421.220	5.260, 2.863	
	SD, SE	3440.7600, 486.5980	8.1569, 1.0040	
	MEDIAN	4470.820	2.243	
	MIN, MAX	44.556, 17099.775	0.570, 46.593	
	5%, 95%	720.720, 10349.600	1.030, 23.940	
	BLQ Values Included	0	34	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	5021.690, 3664.560	6.335, 3.240	
	SD, SE	3238.0900, 462.5840	12.5104, 1.5399	
	MEDIAN	5068.690	2.745	
	MIN, MAX	25.650, 16352.593	0.625, 95.190	
	5%, 95%	735.694, 11711.000	0.915, 17.978	
	BLQ Values Included	0	33	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	5092.020, 3687.210	4.695, 2.968	
	SD, SE	3093.8100, 456.1580	7.0334, 0.8657	
	MEDIAN	5049.490	2.475	
	MIN, MAX	21.379, 13322.126	0.990, 43.940	
	5%, 95%	538.890, 10641.300	1.200, 14.945	
	BLQ Values Included	0	40	
Overall[b]	N	50	66	0.5580
	MEAN, GEO MEAN	4922.140, 3635.760	5.430, 3.243	
	SD, SE	3018.0300, 426.8140	7.9536, 0.9790	
	MEDIAN	5080.990	2.645	
	MIN, MAX	30.528, 13142.500	0.967, 48.539	
	5%, 95%	834.401, 10962.600	1.181, 21.198	

[a] P-value from wilcoxon rank-sum test.

[b] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

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TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Total Cotinine (nmol/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	27684.800, 19415.400	29.851, 16.247	
	SD, SE	19526.3000, 2761.4400	46.2905, 5.6980	
	MEDIAN	25371.900	12.727	
	MIN, MAX	252.855, 97041.223	3.235, 264.415	
	5%, 95%	4090.090, 58734.100	5.845, 135.860	
	BLQ Values Included	0	34	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	28498.100, 20796.400	35.952, 18.386	
	SD, SE	18376.2000, 2625.1700	70.9965, 8.7391	
	MEDIAN	28764.800	15.578	
	MIN, MAX	145.564, 92800.965	3.547, 540.203	
	5%, 95%	4175.060, 66459.800	5.193, 102.025	
	BLQ Values Included	0	33	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	28897.200, 20924.900	26.643, 16.843	
	SD, SE	17557.4000, 2588.7000	39.9143, 4.9131	
	MEDIAN	28655.900	14.046	
	MIN, MAX	121.326, 75603.065	5.618, 249.360	
	5%, 95%	3058.200, 60389.500	6.810, 84.813	
	BLQ Values Included	0	40	
Overall[b]	N	50	66	0.5580
	MEAN, GEO MEAN	27933.100, 20632.900	30.816, 18.402	
	SD, SE	17127.3000, 2422.1700	45.1369, 5.5560	
	MEDIAN	28834.600	15.009	
	MIN, MAX	173.248, 74583.600	5.486, 275.461	
	5%, 95%	4735.230, 62212.600	6.704, 120.299	

[a] P-value from wilcoxon rank-sum test.
[b] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 11 of ?)

Biomarker: Total Cotinine (ug/mg Creatinin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value*
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	3.957, 2.835	0.004, NA	
	SD, SE	2.4437, 0.3456	0.0056, 0.0007	
	MEDIAN	3.861	0.002	
	MIN, MAX	0.042, 10.371	0.000, 0.033	
	5%, 95%	0.473, 8.403	0.001, 0.015	
	BLQ Values Included	0	34	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	3.842, 2.829	0.005, NA	
	SD, SE	2.5305, 0.3615	0.0113, 0.0014	
	MEDIAN	3.699	0.002	
	MIN, MAX	0.029, 12.831	0.000, 0.091	
	5%, 95%	0.636, 8.239	0.001, 0.014	
	BLQ Values Included	0	33	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	4.042, 2.957	0.003, 0.002	
	SD, SE	2.6149, 0.3855	0.0044, 0.0005	
	MEDIAN	3.681	0.002	
	MIN, MAX	0.019, 11.859	0.001, 0.025	
	5%, 95%	0.832, 9.415	0.001, 0.008	
	BLQ Values Included	0	40	
Overall*	N	50	66	0.1154
	MEAN, GEO MEAN	3.950, 2.935	0.004, 0.002	
	SD, SE	2.4396, 0.3450	0.0060, 0.0007	
	MEDIAN	3.804	0.002	
	MIN, MAX	0.030, 10.694	0.001, 0.042	
	5%, 95%	0.733, 8.810	0.001, 0.015	

* P-value from wilcoxon rank-sum test.
* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Total Cotinine (nmol/mg Creatin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	22.456, 16.086	0.021, NA	
	SD, SE	13.8680, 1.9612	0.0319, 0.0039	
	MEDIAN	21.912	0.011	
	MIN, MAX	0.238, 58.855	0.000, 0.187	
	5%, 95%	2.684, 47.687	0.006, 0.085	
	BLQ Values Included	0	34	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	21.801, 16.055	0.028, NA	
	SD, SE	14.3604, 2.0515	0.0642, 0.0079	
	MEDIAN	20.992	0.011	
	MIN, MAX	0.165, 72.816	0.000, 0.516	
	5%, 95%	3.609, 46.756	0.006, 0.079	
	BLQ Values Included	0	33	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	22.940, 16.783	0.019, 0.013	
	SD, SE	14.8394, 2.1880	0.0250, 0.0031	
	MEDIAN	20.887	0.011	
	MIN, MAX	0.108, 67.300	0.006, 0.142	
	5%, 95%	4.722, 53.430	0.006, 0.045	
	BLQ Values Included	0	40	
Overall[*]	N	50	66	0.1222
	MEAN, GEO MEAN	22.417, 16.655	0.023, 0.014	
	SD, SE	13.8448, 1.9579	0.0344, 0.0042	
	MEDIAN	21.585	0.011	
	MIN, MAX	0.170, 60.690	0.004, 0.240	
	5%, 95%	4.162, 49.999	0.006, 0.085	

[a] P-value from wilcoxon rank-sum test.

[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

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TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Free Cotinine (ug/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	1779.840, 1321.430	3.138, 2.091	
	SD, SE	1152.0100, 162.9180	4.4120, 0.5431	
	MEDIAN	1795.820	1.915	
	MIN, MAX	26.790, 6435.660	0.530, 27.900	
	5%, 95%	308.979, 3580.190	0.780, 12.160	
	BLQ Values Included	0	50	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	1817.750, 1365.990	3.672, 2.295	
	SD, SE	1072.6600, 153.2370	7.3843, 0.9089	
	MEDIAN	1731.970	2.180	
	MIN, MAX	13.950, 4366.890	0.625, 59.280	
	5%, 95%	226.118, 3766.020	0.815, 9.804	
	BLQ Values Included	0	48	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	1953.820, 1428.930	2.968, 2.241	
	SD, SE	1298.5800, 191.4640	3.6635, 0.4509	
	MEDIAN	1633.050	2.050	
	MIN, MAX	6.393, 5162.740	0.750, 25.870	
	5%, 95%	354.936, 4856.040	1.068, 7.188	
	BLQ Values Included	0	51	
Overall[b]	N	50	66	0.4576
	MEAN, GEO MEAN	1822.900, 1402.090	3.259, 2.313	
	SD, SE	1058.1900, 149.6500	4.3883, 0.5402	
	MEDIAN	1695.440	2.147	
	MIN, MAX	15.711, 4483.280	0.787, 30.043	
	5%, 95%	374.266, 3774.690	1.092, 12.202	

[a] P-value from wilcoxon rank-sum test.
[b] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 14 of 7)

Biomarker: Free Cotinine (nmol/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	10100.600, 7499.140	17.809, 11.869	
	SD, SE	6537.6300, 924.5610	25.0383, 3.0820	
	MEDIAN	10191.300	10.868	
	MIN, MAX	152.033, 36522.371	3.008, 158.333	
	5%, 95%	1753.460, 20317.600	4.427, 69.008	
	BLQ Values Included	0	50	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	10315.700, 7751.970	20.841, 13.025	
	SD, SE	6087.3400, 869.6210	41.9057, 5.1582	
	MEDIAN	9828.950	12.372	
	MIN, MAX	79.166, 24782.101	3.547, 336.414	
	5%, 95%	1283.220, 21372.200	4.625, 55.638	
	BLQ Values Included	0	48	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	11087.900, 8109.170	16.841, 12.719	
	SD, SE	7369.4100, 1086.5600	20.7903, 2.5591	
	MEDIAN	9267.540	11.634	
	MIN, MAX	36.280, 29298.550	4.256, 146.812	
	5%, 95%	2014.260, 27558.000	6.061, 40.792	
	BLQ Values Included	0	51	
Overall[b]	N	50	66	0.4576
	MEAN, GEO MEAN	10344.900, 7956.860	18.497, 13.128	
	SD, SE	6005.2200, 849.2660	24.9037, 3.0654	
	MEDIAN	9621.610	12.182	
	MIN, MAX	89.160, 25442.600	4.464, 170.496	
	5%, 95%	2123.960, 21421.400	6.195, 69.245	

[a] P-value from wilcoxon rank-sum test.
[b] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminating all BLQ values for the overall statistics.

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Biomarker: Free Cotinine (ug/mg Creatinin)

Timepoint	Statistic	Smokers	Non-Smokers	P value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	1.607, 1.095	0.002, NA	
	SD, SE	1.2214, 0.1727	0.0030, 0.0004	
	MEDIAN	1.401	0.002	
	MIN, MAX	0.025, 5.808	0.000, 0.020	
	5%, 95%	0.203, 3.903	0.000, 0.009	
	BLQ Values Included	0	50	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	1.474, 1.055	0.003, NA	
	SD, SE	1.0867, 0.1552	0.0070, 0.0009	
	MEDIAN	1.263	0.002	
	MIN, MAX	0.016, 5.177	0.000, 0.057	
	5%, 95%	0.148, 3.337	0.000, 0.007	
	BLQ Values Included	0	48	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	1.609, 1.147	0.002, NA	
	SD, SE	1.1859, 0.1749	0.0022, 0.0003	
	MEDIAN	1.333	0.002	
	MIN, MAX	0.006, 5.186	0.000, 0.014	
	5%, 95%	0.383, 4.005	0.001, 0.005	
	BLQ Values Included	0	51	
Overall[b]	N	50	66	0.0872
	MEAN, GEO MEAN	1.575, 1.138	0.002, NA	
	SD, SE	1.1231, 0.1588	0.0034, 0.0004	
	MEDIAN	1.305	0.002	
	MIN, MAX	0.016, 4.572	0.000, 0.026	
	5%, 95%	0.266, 3.886	0.001, 0.007	

[a] P-value from wilcoxon rank-sum test.
[b] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Free Cotinine (nmol/mg Creatin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[4]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	9.121, 6.214	0.013, NA	
	SD, SE	6.9314, 0.9802	0.0172, 0.0021	
	MEDIAN	7.951	0.009	
	MIN, MAX	0.142, 32.960	0.000, 0.114	
	5%, 95%	1.152, 22.150	0.000, 0.051	
	BLQ Values Included	0	50	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	8.365, 5.987	0.017, NA	
	SD, SE	6.1668, 0.8810	0.0394, 0.0048	
	MEDIAN	7.168	0.011	
	MIN, MAX	0.091, 29.379	0.000, 0.323	
	5%, 95%	0.840, 18.937	0.000, 0.040	
	BLQ Values Included	0	48	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	9.133, 6.511	0.012, NA	
	SD, SE	6.7302, 0.9923	0.0124, 0.0015	
	MEDIAN	7.565	0.011	
	MIN, MAX	0.034, 29.431	0.000, 0.079	
	5%, 95%	2.174, 22.728	0.006, 0.028	
	BLQ Values Included	0	51	
Overall*	N	50	66	0.0970
	MEAN, GEO MEAN	8.937, 6.454	0.014, NA	
	SD, SE	6.3738, 0.9014	0.0195, 0.0024	
	MEDIAN	7.403	0.009	
	MIN, MAX	0.089, 25.944	0.000, 0.149	
	5%, 95%	1.510, 22.053	0.004, 0.038	

[4] P-value from wilcoxon rank-sum test.
* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Total 3-hydroxycotinine (ug/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	7831.130, 5355.820	8.239, 3.926	
	SD, SE	6001.9500, 848.8040	14.3955, 1.7720	
	MEDIAN	7518.910	2.948	
	MIN, MAX	80.370, 31233.615	0.800, 84.568	
	5%, 95%	795.900, 18631.300	1.140, 40.870	
	BLQ Values Included	0	24	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	8569.600, 6203.500	11.439, 4.758	
	SD, SE	5412.4100, 773.2020	25.8488, 3.1818	
	MEDIAN	7610.060	3.038	
	MIN, MAX	48.420, 22147.192	1.000, 191.422	
	5%, 95%	842.508, 19677.600	1.148, 43.240	
	BLQ Values Included	0	16	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	8483.710, 5938.860	7.913, 4.073	
	SD, SE	6120.1700, 902.3690	14.4330, 1.7766	
	MEDIAN	6586.050	3.210	
	MIN, MAX	52.295, 27454.900	1.068, 89.830	
	5%, 95%	1258.180, 18489.900	1.350, 44.835	
	BLQ Values Included	0	26	
Overall[b]	N	50	66	0.9843
	MEAN, GEO MEAN	8179.590, 5975.970	9.197, 4.643	
	SD, SE	5444.3900, 769.9540	15.9724, 1.9661	
	MEDIAN	6813.150	3.307	
	MIN, MAX	60.362, 23472.700	1.067, 97.065	
	5%, 95%	827.503, 20711.600	1.355, 42.982	

[a] P-value from wilcoxon rank-sum test.
[b] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

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TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Total 3-hydroxycotinine (nmol/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	40745.400, 27866.300	42.869, 20.426	
	SD, SE	31228.2000, 4416.3300	74.8998, 9.2195	
	MEDIAN	39120.900	15.336	
	MIN, MAX	418.165, 162508.499	4.162, 440.007	
	5%, 95%	4141.070, 96938.400	5.931, 212.647	
	BLQ Values Included	0	24	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	44587.600, 32276.800	59.515, 24.754	
	SD, SE	28160.8000, 4022.9700	134.4910, 16.5547	
	MEDIAN	39595.200	15.804	
	MIN, MAX	251.929, 115231.840	5.203, 995.969	
	5%, 95%	4383.570, 102382.000	5.973, 224.978	
	BLQ Values Included	0	16	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	44140.700, 30899.900	41.171, 21.193	
	SD, SE	31843.2000, 4695.0300	75.0951, 9.2436	
	MEDIAN	34267.200	16.699	
	MIN, MAX	272.091, 142847.845	5.557, 467.385	
	5%, 95%	6546.310, 96202.800	7.024, 233.277	
	BLQ Values Included	0	26	
Overall[*]	N	50	66	0.9843
	MEAN, GEO MEAN	42558.500, 31093.000	47.852, 24.159	
	SD, SE	28327.2000, 4006.0800	83.1043, 10.2294	
	MEDIAN	35448.800	17.204	
	MIN, MAX	314.062, 122129.000	5.550, 505.029	
	5%, 95%	4305.500, 107763.000	7.048, 223.634	

[a] P-value from wilcoxon rank-sum test.
[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Total 3-hydroxycotinine (ug/mg Creatinin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	6.449, 4.438	0.006, NA	
	SD, SE	4.6629, 0.6594	0.0097, 0.0012	
	MEDIAN	5.375	0.002	
	MIN, MAX	0.076, 18.383	0.000, 0.052	
	5%, 95%	0.370, 15.997	0.001, 0.022	
	BLQ Values Included	0	24	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	6.772, 4.788	0.008, NA	
	SD, SE	4.9259, 0.7037	0.0155, 0.0019	
	MEDIAN	5.554	0.003	
	MIN, MAX	0.054, 20.024	0.000, 0.099	
	5%, 95%	0.503, 18.134	0.001, 0.027	
	BLQ Values Included	0	16	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	6.924, 4.762	0.005, 0.003	
	SD, SE	5.3191, 0.7843	0.0079, 0.0010	
	MEDIAN	5.015	0.002	
	MIN, MAX	0.046, 23.294	0.001, 0.050	
	5%, 95%	1.148, 17.524	0.001, 0.021	
	BLQ Values Included	0	26	
Overall[b]	N	50	66	0.3418
	MEAN, GEO MEAN	6.692, 4.787	0.006, 0.003	
	SD, SE	4.8008, 0.6789	0.0094, 0.0012	
	MEDIAN	5.297	0.003	
	MIN, MAX	0.059, 20.195	0.001, 0.046	
	5%, 95%	0.653, 16.414	0.001, 0.026	

[a] P-value from wilcoxon rank-sum test.
[b] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminating all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL.
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Biomarker: Total 3-hydroxycotinine (nmol/mg Creatin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value*
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	33.556, 23.089	0.030, NA	
	SD, SE	24.2611, 3.4310	0.0506, 0.0062	
	MEDIAN	27.964	0.010	
	MIN, MAX	0.395, 95.647	0.000, 0.271	
	5%, 95%	1.925, 83.232	0.005, 0.114	
	BLQ Values Included	0	24	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	35.235, 24.911	0.040, NA	
	SD, SE	25.6293, 3.6613	0.0805, 0.0099	
	MEDIAN	28.897	0.013	
	MIN, MAX	0.281, 104.185	0.000, 0.515	
	5%, 95%	2.617, 94.351	0.005, 0.140	
	BLQ Values Included	0	16	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	36.028, 24.775	0.026, 0.015	
	SD, SE	27.6752, 4.0805	0.0412, 0.0051	
	MEDIAN	26.091	0.010	
	MIN, MAX	0.239, 121.199	0.005, 0.260	
	5%, 95%	5.973, 91.177	0.005, 0.109	
	BLQ Values Included	0	26	
Overall*	N	50	66	0.3433
	MEAN, GEO MEAN	34.818, 24.906	0.032, 0.017	
	SD, SE	24.9784, 3.5325	0.0490, 0.0060	
	MEDIAN	27.557	0.016	
	MIN, MAX	0.305, 105.075	0.003, 0.241	
	5%, 95%	3.399, 85.402	0.005, 0.135	

* P-value from wilcoxon rank-sum test.
* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminating all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Free 3-hydroxycotinine (ug/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	5745.430, 3951.120	6.880, 3.530	
	SD, SE	4337.8600, 613.4660	11.5292, 1.4191	
	MEDIAN	5379.730	2.763	
	MIN, MAX	42.441, 20367.215	0.855, 67.936	
	5%, 95%	645.000, 15778.200	1.030, 30.820	
	BLQ Values Included	0	23	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	6615.880, 4719.000	9.154, 4.133	
	SD, SE	4263.6400, 609.0920	20.5086, 2.5244	
	MEDIAN	6368.090	2.918	
	MIN, MAX	24.210, 18154.368	0.750, 152.550	
	5%, 95%	722.866, 15032.800	1.023, 31.960	
	BLQ Values Included	0	18	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	6170.890, 4344.160	6.458, 3.634	
	SD, SE	4218.9800, 622.0550	11.1448, 1.3718	
	MEDIAN	5068.950	2.894	
	MIN, MAX	21.379, 15121.480	0.990, 72.800	
	5%, 95%	976.686, 14222.600	1.300, 31.610	
	BLQ Values Included	0	31	
Overall[*]	N	50	66	0.9308
	MEAN, GEO MEAN	6093.850, 4444.840	7.498, 4.099	
	SD, SE	3968.4900, 561.2300	12.6611, 1.5585	
	MEDIAN	5310.140	3.179	
	MIN, MAX	29.343, 17222.200	1.038, 76.940	
	5%, 95%	696.229, 12433.200	1.303, 33.167	

[a] P-value from wilcoxon rank-sum test.

[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminating all BLQ values for the overall statistics.

PM300355245

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Free 3-hydroxycotinine (nmol/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	29893.500, 20557.700	35.797, 18.368	
	SD, SE	22569.9000, 3191.8600	59.9865, 7.3838	
	MEDIAN	27990.700	14.373	
	MIN, MAX	220.821, 105970.620	4.449, 353.471	
	5%, 95%	3355.940, 82094.200	5.359, 160.356	
	BLQ Values Included	0	23	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	34422.400, 24553.000	47.631, 21.505	
	SD, SE	22183.7000, 3169.1100	106.7060, 13.1346	
	MEDIAN	33133.100	15.180	
	MIN, MAX	125.965, 94457.177	3.902, 793.718	
	5%, 95%	3761.070, 78215.800	5.323, 166.288	
	BLQ Values Included	0	18	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	32107.100, 22602.700	33.601, 18.909	
	SD, SE	21951.4000, 3236.5500	57.9863, 7.1376	
	MEDIAN	26373.800	15.058	
	MIN, MAX	111.235, 78677.060	5.151, 378.778	
	5%, 95%	5081.700, 74000.000	6.764, 164.467	
	BLQ Values Included	0	31	
Overall*	N	50	66	0.9308
	MEAN, GEO MEAN	31706.300, 23126.500	39.009, 21.327	
	SD, SE	20648.1000, 2920.0800	65.8755, 8.1087	
	MEDIAN	27628.600	16.540	
	MIN, MAX	152.674, 89607.000	5.402, 400.319	
	5%, 95%	3622.480, 64690.100	6.778, 172.570	

[a] P-value from wilcoxon rank-sum test.
* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminating all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Free 3-hydroxycotinine (ug/mg Creatinin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	4.766, 3.274	0.005, NA	
	SD, SE	3.4639, 0.4899	0.0080, 0.0010	
	MEDIAN	4.258	0.002	
	MIN, MAX	0.040, 14.351	0.000, 0.042	
	5%, 95%	0.300, 12.478	0.001, 0.019	
	BLQ Values Included	0	23	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	5.168, 3.642	0.006, NA	
	SD, SE	3.6621, 0.5232	0.0122, 0.0015	
	MEDIAN	4.126	0.002	
	MIN, MAX	0.027, 15.297	0.000, 0.079	
	5%, 95%	0.400, 12.899	0.001, 0.019	
	BLQ Values Included	0	18	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	5.034, 3.484	0.004, 0.003	
	SD, SE	3.6689, 0.5410	0.0062, 0.0008	
	MEDIAN	3.689	0.002	
	MIN, MAX	0.019, 15.542	0.001, 0.041	
	5%, 95%	0.770, 11.642	0.001, 0.017	
	BLQ Values Included	0	31	
Overall[*]	N	50	66	0.2607
	MEAN, GEO MEAN	4.971, 3.566	0.005, 0.003	
	SD, SE	3.4521, 0.4882	0.0074, 0.0009	
	MEDIAN	4.113	0.003	
	MIN, MAX	0.029, 14.264	0.001, 0.037	
	5%, 95%	0.497, 12.705	0.001, 0.019	

[a] P-value from wilcoxon rank-sum test.

[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

PM3003565247

TABLE 14.1.1-2

STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Free 3-hydroxycotinine (nmol/mg Creatin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	24.797, 17.032	0.025, NA	
	SD, SE	18.0226, 2.5488	0.0415, 0.0051	
	MEDIAN	22.155	0.010	
	MIN, MAX	0.208, 74.668	0.000, 0.219	
	5%, 95%	1.561, 64.923	0.005, 0.099	
	BLQ Values Included	0	23	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	26.891, 18.948	0.032, NA	
	SD, SE	19.0536, 2.7219	0.0636, 0.0078	
	MEDIAN	21.468	0.010	
	MIN, MAX	0.140, 79.590	0.000, 0.411	
	5%, 95%	2.081, 67.113	0.005, 0.099	
	BLQ Values Included	0	18	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	26.191, 18.128	0.022, 0.013	
	SD, SE	19.0893, 2.8146	0.0323, 0.0040	
	MEDIAN	19.194	0.010	
	MIN, MAX	0.099, 80.865	0.005, 0.213	
	5%, 95%	4.006, 60.573	0.005, 0.088	
	BLQ Values Included	0	31	
Overall[*]	N	50	66	0.2766
	MEAN, GEO MEAN	25.862, 18.548	0.026, 0.015	
	SD, SE	17.9615, 2.5401	0.0386, 0.0048	
	MEDIAN	21.398	0.013	
	MIN, MAX	0.149, 74.215	0.003, 0.194	
	5%, 95%	2.586, 66.106	0.005, 0.099	

[a] P-value from wilcoxon rank-sum test.
[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 25 of ?)

Biomarker: 3-hydroxycotinine glucuronide (nmol/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	49	54	0.0000
	MEAN, GEO MEAN	11096.200, 6918.750	9.097, NA	
	SD, SE	10472.9000, 1496.1300	17.1446, 2.3331	
	MEDIAN	8366.020	2.082	
	MIN, MAX	197.400, 56537.828	0.000, 86.504	
	5%, 95%	757.770, 26230.400	0.000, 52.260	
	BLQ Values Included	0	23	
Week 2	N	49	60	0.0000
	MEAN, GEO MEAN	10165.200, 7180.450	13.441, NA	
	SD, SE	7498.0900, 1071.1600	30.2054, 3.8995	
	MEDIAN	8992.600	1.533	
	MIN, MAX	126.000, 38811.750	0.000, 202.270	
	5%, 95%	622.537, 24231.800	0.000, 60.436	
	BLQ Values Included	0	18	
Week 3	N	45	62	0.0000
	MEAN, GEO MEAN	12310.200, 8164.710	8.214, NA	
	SD, SE	11374.8000, 1695.6600	18.5892, 2.3608	
	MEDIAN	8468.660	0.995	
	MIN, MAX	160.868, 64170.700	0.000, 88.660	
	5%, 95%	1725.520, 31402.400	0.000, 42.273	
	BLQ Values Included	0	31	
Overall[b]	N	50	65	0.8759
	MEAN, GEO MEAN	10913.700, 7675.780	9.891, NA	
	SD, SE	8872.2500, 1254.7300	19.2777, 2.3911	
	MEDIAN	9018.320	2.395	
	MIN, MAX	161.423, 49042.900	0.000, 104.787	
	5%, 95%	828.268, 27011.000	0.000, 49.847	

[a] P-value from wilcoxon rank-sum test.
[b] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: 3-hydroxycotinine glucuronide (nmol/mg Creatin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	49	54	0.0000
	MEAN, GEO MEAN	8.953, 5.755	0.006, NA	
	SD, SE	7.6799, 1.0971	0.0113, 0.0015	
	MEDIAN	6.754	0.002	
	MIN, MAX	0.182, 37.904	0.000, 0.053	
	5%, 95%	0.365, 24.215	0.000, 0.034	
	BLQ Values Included	0	23	
Week 2	N	49	60	0.0000
	MEAN, GEO MEAN	8.344, 5.542	0.009, NA	
	SD, SE	7.7423, 1.1060	0.0187, 0.0024	
	MEDIAN	5.726	0.001	
	MIN, MAX	0.141, 37.773	0.000, 0.105	
	5%, 95%	0.539, 24.592	0.000, 0.047	
	BLQ Values Included	0	18	
Week 3	N	45	62	0.0000
	MEAN, GEO MEAN	10.066, 6.491	0.005, NA	
	SD, SE	9.7261, 1.4499	0.0101, 0.0013	
	MEDIAN	6.677	0.001	
	MIN, MAX	0.141, 41.198	0.000, 0.050	
	5%, 95%	1.718, 30.778	0.000, 0.030	
	BLQ Values Included	0	31	
Overall[b]	N	50	65	0.7982
	MEAN, GEO MEAN	9.000, 6.118	0.006, NA	
	SD, SE	7.9341, 1.1220	0.0115, 0.0014	
	MEDIAN	6.640	0.002	
	MIN, MAX	0.156, 38.958	0.000, 0.052	
	5%, 95%	0.813, 26.074	0.000, 0.036	

[a] P-value from wilcoxon rank-sum test.

[b] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: 3-hydroxycotinine glucuronide (nmol/L)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	49	54	0.0000
	MEAN, GEO MEAN	6895.8, 4024.9	5.9, NA	
	SD, SE	7654.61, 1093.52	14.15, 1.93	
	MEDIAN	5136.9	1.9	
	MIN, MAX	79.6, 48117.3	0.0, 98.3	
	5%, 95%	649.9, 16292.2	0.0, 21.9	
	BLQ Values Included	0	23	
Week 2	N	49	60	0.0000
	MEAN, GEO MEAN	6711.8, 4641.4	8.7, NA	
	SD, SE	5108.69, 729.81	24.88, 3.21	
	MEDIAN	5545.9	1.6	
	MIN, MAX	140.0, 23215.8	0.0, 179.0	
	5%, 95%	596.3, 15749.0	0.0, 36.5	
	BLQ Values Included	0	18	
Week 3	N	45	62	0.0000
	MEAN, GEO MEAN	7528.7, 4762.9	4.5, NA	
	SD, SE	8062.18, 1201.84	10.28, 1.31	
	MEDIAN	5225.9	0.5	
	MIN, MAX	153.5, 40612.0	0.0, 68.2	
	5%, 95%	927.7, 29168.5	0.0, 24.5	
	BLQ Values Included	0	31	
Overall[*]	N	50	65	0.6395
	MEAN, GEO MEAN	6902.1, 4770.7	6.3, NA	
	SD, SE	5566.90, 787.28	14.01, 1.74	
	MEDIAN	5704.9	1.6	
	MIN, MAX	144.5, 25999.9	0.0, 83.3	
	5%, 95%	640.7, 19989.6	0.0, 22.2	

[a] P-value from wilcoxon rank-sum test.
[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

PM3003565251

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Total NNAL (pg/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	49	66	0.0000
	MEAN, GEO MEAN	860759.000, 585425.000	14427.700, 10836.700	
	SD, SE	613472.0000, 87638.9000	21975.5000, 2704.9900	
	MEDIAN	785030.000	9870.000	
	MIN, MAX	8460.000, 2307613.000	3180.000, 181986.000	
	5%, 95%	37125.000, 2084330.000	4680.000, 21000.000	
	BLQ Values Included	1	59	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	914458.000, 627899.000	14417.800, 12348.400	
	SD, SE	709365.0000, 104590.0000	10349.1000, 1273.8800	
	MEDIAN	783064.000	12000.000	
	MIN, MAX	6288.000, 4046406.000	4500.000, 61490.000	
	5%, 95%	97520.000, 1964160.000	6408.000, 28440.000	
	BLQ Values Included	1	59	
Overall*	N	50	66	0.1165
	MEAN, GEO MEAN	874402.000, 615372.000	14422.800, 11984.200	
	SD, SE	592616.0000, 83808.5000	12891.4000, 1586.8300	
	MEDIAN	839770.000	11130.000	
	MIN, MAX	7374.000, 2468210.000	3840.000, 97218.000	
	5%, 95%	66085.000, 2080140.000	6400.000, 30583.000	

[a] P-value from wilcoxon rank-sum test.
* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 29 of ?)

Biomarker: Total NNAL (pmol/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	49	66	0.0000
	MEAN, GEO MEAN	4114.430, 2798.330	68.965, 51.799	
	SD, SE	2932.4000, 418.9140	105.0430, 12.9299	
	MEDIAN	3752.440	47.179	
	MIN, MAX	40.439, 11030.390	15.200, 869.893	
	5%, 95%	177.458, 9963.110	22.370, 100.380	
	BLQ Values Included	1	59	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	4371.110, 3001.360	68.917, 59.025	
	SD, SE	3390.7700, 499.9410	49.4686, 6.0892	
	MEDIAN	3743.050	57.360	
	MIN, MAX	30.057, 19341.821	21.510, 293.922	
	5%, 95%	466.146, 9388.690	30.630, 135.943	
	BLQ Values Included	1	59	
Overall[*]	N	50	66	0.1159
	MEAN, GEO MEAN	4179.640, 2941.480	68.941, 57.285	
	SD, SE	2832.7000, 400.6040	61.6210, 7.5850	
	MEDIAN	4014.100	53.202	
	MIN, MAX	35.248, 11798.000	18.355, 464.702	
	5%, 95%	315.887, 9943.070	30.592, 146.187	

[a] P-value from wilcoxon rank-sum test.
[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminating all BLQ values for the overall statistics.

PM3003565253

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Total NNAL (pg/mg Creatinin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value*
Week 1	N	49	66	0.0000
	MEAN, GEO MEAN	738.054, 487.673	10.548, 7.669	
	SD, SE	585.2480, 83.6069	16.2628, 2.0018	
	MEDIAN	625.442	9.003	
	MIN, MAX	7.958, 2575.145	1.932, 134.956	
	5%, 95%	30.892, 2010.240	2.683, 17.699	
	BLQ Values Included	1	59	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	721.933, 503.456	11.157, 8.773	
	SD, SE	524.1050, 77.2750	9.2533, 1.1390	
	MEDIAN	580.019	8.596	
	MIN, MAX	5.525, 2275.640	2.158, 53.767	
	5%, 95%	65.031, 2004.780	2.923, 30.000	
	BLQ Values Included	1	59	
Overall*	N	50	66	0.0039
	MEAN, GEO MEAN	726.768, 500.586	10.853, 8.600	
	SD, SE	532.2830, 75.2762	9.7040, 1.1945	
	MEDIAN	617.106	8.560	
	MIN, MAX	6.742, 2250.750	2.131, 70.371	
	5%, 95%	43.687, 1877.500	3.508, 27.562	

* P-value from wilcoxon rank-sum test.
* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 31 of ?)

Biomarker: Total NNAL (pmol/mg Creatin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	49	66	0.0000
	MEAN, GEO MEAN	3.528, 2.331	0.050, 0.037	
	SD, SE	2.7975, 0.3996	0.0777, 0.0096	
	MEDIAN	2.990	0.043	
	MIN, MAX	0.038, 12.309	0.009, 0.645	
	5%, 95%	0.148, 9.609	0.013, 0.085	
	BLQ Values Included	1	59	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	3.451, 2.406	0.053, 0.042	
	SD, SE	2.5053, 0.3694	0.0442, 0.0054	
	MEDIAN	2.773	0.041	
	MIN, MAX	0.026, 10.878	0.010, 0.257	
	5%, 95%	0.311, 9.583	0.014, 0.143	
	BLQ Values Included	1	59	
Overall[*]	N	50	66	0.0039
	MEAN, GEO MEAN	3.474, 2.393	0.052, 0.041	
	SD, SE	2.5443, 0.3598	0.0464, 0.0057	
	MEDIAN	2.950	0.042	
	MIN, MAX	0.032, 10.759	0.010, 0.337	
	5%, 95%	0.209, 8.975	0.017, 0.132	

[a] P-value from wilcoxon rank-sum test.
[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 32 of ?)

Biomarker: Free NNAL (pg/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	249288.000, 181853.000	11517.800, 10146.400	
	SD, SE	180187.0000, 25482.3000	6048.7800, 744.5530	
	MEDIAN	224364.000	9240.000	
	MIN, MAX	8460.000, 852355.000	3180.000, 30796.800	
	5%, 95%	23940.000, 649460.000	4800.000, 22800.000	
	BLQ Values Included	2	62	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	278384.000, 210748.000	14085.400, 12211.000	
	SD, SE	174392.0000, 25712.7000	9300.8400, 1144.8500	
	MEDIAN	241481.000	12090.000	
	MIN, MAX	6288.000, 781704.000	4500.000, 57240.000	
	5%, 95%	52440.000, 558700.000	5940.000, 28440.000	
	BLQ Values Included	1	62	
Overall*	N	50	66	0.1077
	MEAN, GEO MEAN	259455.000, 199547.000	12801.600, 11589.200	
	SD, SE	160409.0000, 22685.3000	6057.8600, 745.6710	
	MEDIAN	239327.000	11217.000	
	MIN, MAX	7374.000, 589956.000	3840.000, 32310.000	
	5%, 95%	47776.000, 584079.000	6450.000, 24998.400	

[a] P-value from wilcoxon rank-sum test.
* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 33 of ?)

Biomarker: Free NNAL (pmol/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value*
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	1191.600, 869.259	55.055, 48.500	
	SD, SE	861.2930, 121.8050	28.9132, 3.5590	
	MEDIAN	1072.460	44.168	
	MIN, MAX	40.439, 4074.257	15.200, 147.209	
	5%, 95%	114.433, 3104.420	22.944, 108.984	
	BLQ Values Included	2	62	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	1330.680, 1007.380	67.328, 58.369	
	SD, SE	833.5930, 122.9070	44.4580, 5.4724	
	MEDIAN	1154.280	57.790	
	MIN, MAX	30.057, 3736.545	21.510, 273.607	
	5%, 95%	250.663, 2670.590	28.393, 135.943	
	BLQ Values Included	1	62	
Overall⁺	N	50	66	0.1070
	MEAN, GEO MEAN	1240.200, 953.832	61.192, 55.397	
	SD, SE	766.7570, 108.4360	28.9566, 3.5643	
	MEDIAN	1143.980	53.618	
	MIN, MAX	35.248, 2819.990	18.355, 154.442	
	5%, 95%	228.369, 2791.900	30.831, 119.493	

* P-value from wilcoxon rank-sum test.
⁺ Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1,234' will be used as '1,234') for the averages across visits within a subject thus eliminating all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 34 of ?)

Biomarker: Free NNAL (pg/mg Creatinin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	219.163, 150.676	8.520, 7.180	
	SD, SE	181.7280, 25.7002	5.1180, 0.6300	
	MEDIAN	178.751	8.222	
	MIN, MAX	7.958, 813.359	1.932, 29.814	
	5%, 95%	13.730, 612.755	2.911, 17.143	
	BLQ Values Included	2	62	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	226.894, 168.981	10.544, 8.676	
	SD, SE	159.6470, 23.5386	6.8346, 0.8413	
	MEDIAN	194.555	9.325	
	MIN, MAX	5.525, 724.324	2.158, 30.264	
	5%, 95%	47.240, 547.012	2.923, 29.412	
	BLQ Values Included	1	62	
Overall*	N	50	66	0.0016
	MEAN, GEO MEAN	223.061, 162.757	9.532, 8.256	
	SD, SE	167.5170, 23.6904	5.0078, 0.6164	
	MEDIAN	185.127	8.678	
	MIN, MAX	6.742, 768.842	2.364, 23.043	
	5%, 95%	34.373, 535.268	3.185, 19.673	

[a] P-value from wilcoxon rank-sum test.
* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 35 of ?)

Biomarker: Free NNAL (pmol/mg Creatin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	1.048, 0.720	0.041, 0.034	
	SD, SE	0.8687, 0.1229	0.0245, 0.0030	
	MEDIAN	0.855	0.039	
	MIN, MAX	0.038, 3.888	0.009, 0.143	
	5%, 95%	0.066, 2.929	0.014, 0.082	
	BLQ Values Included	2	62	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	1.085, 0.807	0.050, 0.041	
	SD, SE	0.7631, 0.1125	0.0326, 0.0040	
	MEDIAN	0.930	0.045	
	MIN, MAX	0.026, 3.462	0.010, 0.145	
	5%, 95%	0.226, 2.615	0.014, 0.141	
	BLQ Values Included	1	62	
Overall*	N	50	66	0.0017
	MEAN, GEO MEAN	1.066, 0.778	0.046, 0.040	
	SD, SE	0.8008, 0.1132	0.0239, 0.0029	
	MEDIAN	0.885	0.042	
	MIN, MAX	0.032, 3.675	0.011, 0.110	
	5%, 95%	0.164, 2.559	0.015, 0.094	

[a] P-value from wilcoxon rank-sum test.
* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

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TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: NNAL-Glucuronide (pmol/L)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	49	62	0.0000
	MEAN, GEO MEAN	1889.1, NA	17.8, NA	
	SD, SE	1803.06, 257.58	111.46, 14.16	
	MEDIAN	1564.5	0.0	
	MIN, MAX	0.0, 7462.5	0.0, 859.0	
	5%, 95%	35.9, 5782.8	0.0, 13.4	
	BLQ Values Included	2	62	
Week 3	N	46	64	0.0000
	MEAN, GEO MEAN	1872.0, NA	3.7, NA	
	SD, SE	1907.93, 281.31	15.05, 1.88	
	MEDIAN	1352.5	0.0	
	MIN, MAX	0.0, 9536.6	0.0, 99.9	
	5%, 95%	35.4, 5911.9	0.0, 28.2	
	BLQ Values Included	1	62	
Overall[*]	N	50	66	0.5865
	MEAN, GEO MEAN	1859.5, NA	10.2, NA	
	SD, SE	1623.98, 229.67	54.69, 6.73	
	MEDIAN	1521.7	0.0	
	MIN, MAX	0.0, 5778.3	0.0, 429.5	
	5%, 95%	29.9, 5668.4	0.0, 28.0	

[a] P-value from wilcoxon rank-sum test.
[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminating all BLQ values for the overall statistics.

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TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 37 of ?)

Biomarker: NNAL-Glucuronide (pmol/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	49	62	0.0000
	MEAN, GEO MEAN	2908.930, NA	17.964, NA	
	SD, SE	2273.5200, 324.7890	108.8050, 13.8183	
	MEDIAN	2416.280	0.000	
	MIN, MAX	0.000, 8788.668	0.000, 841.820	
	5%, 95%	98.725, 7630.840	0.000, 21.417	
	BLQ Values Included	2	62	
Week 3	N	46	64	0.0000
	MEAN, GEO MEAN	3040.440, NA	7.025, NA	
	SD, SE	2849.3100, 420.1080	29.5386, 3.6923	
	MEDIAN	2468.520	0.000	
	MIN, MAX	0.000, 17356.612	0.000, 168.831	
	5%, 95%	65.136, 6821.100	0.000, 36.120	
	BLQ Values Included	1	62	
Overall*	N	50	66	0.5845
	MEAN, GEO MEAN	2940.090, NA	11.844, NA	
	SD, SE	2262.4900, 319.9650	54.6839, 6.7311	
	MEDIAN	2635.940	0.000	
	MIN, MAX	0.000, 10073.400	0.000, 420.910	
	5%, 95%	59.593, 7804.880	0.000, 80.217	

[a] P-value from wilcoxon rank-sum test.
* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminating all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 38 of ?)

Biomarker: NNAL-Glucuronide (pmol/mg Creatin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	49	62	0.0000
	MEAN, GEO MEAN	2.466, NA	0.013, NA	
	SD, SE	2.0265, 0.2895	0.0801, 0.0102	
	MEDIAN	2.190	0.000	
	MIN, MAX	0.000, 8.643	0.000, 0.624	
	5%, 95%	0.082, 7.656	0.000, 0.009	
	BLQ Values Included	2	62	
Week 3	N	46	64	0.0000
	MEAN, GEO MEAN	2.366, NA	0.006, NA	
	SD, SE	1.9608, 0.2891	0.0269, 0.0034	
	MEDIAN	1.823	0.000	
	MIN, MAX	0.000, 9.761	0.000, 0.171	
	5%, 95%	0.043, 6.216	0.000, 0.019	
	BLQ Values Included	1	62	
Overall[*]	N	50	66	0.4528
	MEAN, GEO MEAN	2.409, NA	0.009, NA	
	SD, SE	1.8649, 0.2637	0.0409, 0.0050	
	MEDIAN	1.986	0.000	
	MIN, MAX	0.000, 7.312	0.000, 0.312	
	5%, 95%	0.051, 6.478	0.000, 0.065	

[a] P-value from wilcoxon rank-sum test.
[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminating all BLQ values for the overall statistics.

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TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 39 of ?)

Biomarker: Total Nicotine (nmol/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	3	0.0282
	MEAN, GEO MEAN	13340.100, 10943.400	31.361, 28.707	
	SD, SE	8949.6800, 3653.6900	14.0233, 8.0963	
	MEDIAN	10205.000	38.100	
	MIN, MAX	5497.128, 25327.788	15.240, 40.742	
	5%, 95%	5497.130, 25327.800	15.240, 40.742	
	BLQ Values Included	0	3	
Week 2	N	6	5	0.0081
	MEAN, GEO MEAN	12369.000, 11224.600	40.565, 40.139	
	SD, SE	5727.7200, 2338.3300	6.8172, 3.0487	
	MEDIAN	11942.600	36.830	
	MIN, MAX	6300.791, 19866.660	34.925, 51.376	
	5%, 95%	6300.790, 19866.700	34.925, 51.376	
	BLQ Values Included	0	4	
Week 3	N	5	7	0.0058
	MEAN, GEO MEAN	9383.020, 8059.170	29.054, 27.748	
	SD, SE	4868.4500, 2177.2400	9.1357, 3.4530	
	MEDIAN	9143.680	26.213	
	MIN, MAX	2723.706, 14156.770	16.002, 40.640	
	5%, 95%	2723.710, 14156.800	16.002, 40.640	
	BLQ Values Included	0	7	
Overall[b]	N	6	7	0.8303
	MEAN, GEO MEAN	12229.600, 10705.600	31.847, 31.289	
	SD, SE	6469.9800, 2641.3600	6.2904, 2.3775	
	MEDIAN	11297.300	33.528	
	MIN, MAX	4876.530, 19783.700	22.860, 39.298	
	5%, 95%	4876.530, 19783.700	22.860, 39.298	

[a] P-value from wilcoxon rank-sum test.
[b] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 40 of ?)

Biomarker: Total Nicotine (ug/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	3	0.0282
	MEAN, GEO MEAN	2163.760, 1775.030	5.087, 4.656	
	SD, SE	1451.6400, 592.6290	2.2745, 1.3132	
	MEDIAN	1655.260	6.180	
	MIN, MAX	891.634, 4108.167	2.472, 6.608	
	5%, 95%	891.634, 4108.170	2.472, 6.608	
	BLQ Values Included	0	3	
Week 2	N	6	5	0.0081
	MEAN, GEO MEAN	2006.250, 1820.620	6.580, 6.511	
	SD, SE	929.0360, 379.2770	1.1057, 0.4945	
	MEDIAN	1937.100	5.974	
	MIN, MAX	1021.988, 3222.372	5.665, 8.333	
	5%, 95%	1021.990, 3222.370	5.665, 8.333	
	BLQ Values Included	0	4	
Week 3	N	5	7	0.0058
	MEAN, GEO MEAN	1521.930, 1307.200	4.713, 4.501	
	SD, SE	789.6630, 353.1480	1.4816, 0.5600	
	MEDIAN	1483.110	4.252	
	MIN, MAX	441.785, 2296.228	2.596, 6.592	
	5%, 95%	441.785, 2296.230	2.596, 6.592	
	BLQ Values Included	0	7	
Overall[*]	N	6	7	0.8303
	MEAN, GEO MEAN	1983.640, 1736.450	5.166, 5.075	
	SD, SE	1049.4300, 428.4280	1.0202, 0.3856	
	MEDIAN	1832.430	5.439	
	MIN, MAX	790.974, 3208.920	3.708, 6.374	
	5%, 95%	790.974, 3208.920	3.708, 6.374	

[a] P-value from wilcoxon rank-sum test.

[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 41 of 7)

Biomarker: Free Nicotine (nmol/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value*
Week 1	N	6	3	0.0282
	MEAN, GEO MEAN	9618.830, 7479.920	7.072, 6.609	
	SD, SE	7409.9900, 3025.1100	3.0623, 1.7680	
	MEDIAN	6290.340	6.936	
	MIN, MAX	3259.536, 19895.897	4.080, 10.200	
	5%, 95%	3259.540, 19895.900	4.080, 10.200	
	BLQ Values Included	0	3	
Week 2	N	6	2	0.0668
	MEAN, GEO MEAN	8208.240, 7076.090	7.548, 6.393	
	SD, SE	4490.9800, 1833.4400	5.6738, 4.0120	
	MEDIAN	8354.880	7.548	
	MIN, MAX	3484.431, 13388.466	3.536, 11.560	
	5%, 95%	3484.430, 13388.500	3.536, 11.560	
	BLQ Values Included	0	2	
Week 3	N	5	4	0.0200
	MEAN, GEO MEAN	6005.220, 5280.690	7.076, 6.689	
	SD, SE	2926.7700, 1308.8900	2.7799, 1.3899	
	MEDIAN	7103.310	6.569	
	MIN, MAX	2126.580, 8844.885	4.284, 10.880	
	5%, 95%	2126.580, 8844.890	4.284, 10.880	
	BLQ Values Included	0	4	
Overall*	N	6	6	1.0000
	MEAN, GEO MEAN	8471.510, 7224.180	8.359, 7.845	
	SD, SE	5060.2400, 2065.8400	2.9810, 1.2170	
	MEDIAN	7066.450	8.609	
	MIN, MAX	3555.830, 15498.700	3.967, 11.560	
	5%, 95%	3555.830, 15498.700	3.967, 11.560	

* P-value from wilcoxon rank-sum test.
* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

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TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 42 of ?)

Biomarker: Free Nicotine (ug/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value*
Week 1	N	(6)	3	0.0282
	MEAN, GEO MEAN	1560.170, 1213.240	1.147, 1.072	
	SD, SE	1201.9000, 490.6730	0.4964, 0.2866	
	MEDIAN	1020.290	1.125	
	MIN, MAX	528.697, 3227.114	0.662, 1.654	
	5%, 95%	528.697, 3227.110	0.662, 1.654	
	BLQ Values Included	0	3	
Week 2	N	6	2	0.0668
	MEAN, GEO MEAN	1331.380, 1147.740	1.225, 1.037	
	SD, SE	728.4370, 297.3830	0.9199, 0.6505	
	MEDIAN	1355.160	1.225	
	MIN, MAX	565.175, 2171.609	0.574, 1.875	
	5%, 95%	565.175, 2171.610	0.574, 1.875	
	BLQ Values Included	0	2	
Week 3	N	5	4	0.0200
	MEAN, GEO MEAN	974.047, 856.528	1.148, 1.085	
	SD, SE	474.7230, 212.3020	0.4509, 0.2255	
	MEDIAN	1152.160	1.066	
	MIN, MAX	344.931, 1434.640	0.695, 1.765	
	5%, 95%	344.931, 1434.640	0.695, 1.765	
	BLQ Values Included	0	4	
Overall*	N	6	6	1.0000
	MEAN, GEO MEAN	1374.080, 1171.760	1.356, 1.273	
	SD, SE	820.7720, 335.0790	0.4833, 0.1973	
	MEDIAN	1146.180	1.396	
	MIN, MAX	576.756, 2513.890	0.644, 1.875	
	5%, 95%	576.756, 2513.890	0.644, 1.875	

* P-value from wilcoxon rank-sum test.
* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 43 of ?)

Biomarker: Total Cotinine (nmol/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value*
Week 1	N	6	6	0.0051
	MEAN, GEO MEAN	21538.800, 17732.700	22.473, 20.850	
	SD, SE	11226.9000, 4583.3600	9.6010, 3.9196	
	MEDIAN	21849.600	19.360	
	MIN, MAX	3897.624, 34631.742	13.200, 35.288	
	5%, 95%	3897.620, 34631.700	13.200, 35.288	
	BLQ Values Included	0	6	
Week 2	N	6	7	0.0034
	MEAN, GEO MEAN	21942.800, 19820.900	25.654, 23.904	
	SD, SE	8654.6000, 3533.2300	9.4756, 3.5815	
	MEDIAN	25091.500	30.250	
	MIN, MAX	6895.195, 28995.975	11.440, 37.400	
	5%, 95%	6895.200, 28996.000	11.440, 37.400	
	BLQ Values Included	0	7	
Week 3	N	5	7	0.0058
	MEAN, GEO MEAN	18359.900, 13658.400	22.446, 21.506	
	SD, SE	11454.7000, 5122.7000	7.1197, 2.6910	
	MEDIAN	20410.500	20.680	
	MIN, MAX	2437.770, 31425.320	13.860, 32.010	
	5%, 95%	2437.770, 31425.300	13.860, 32.010	
	BLQ Values Included	0	7	
Overall*	N	6	8	0.8983
	MEAN, GEO MEAN	20878.300, 17708.700	23.599, 22.442	
	SD, SE	9961.1900, 4066.6400	7.6547, 2.7064	
	MEDIAN	22902.100	23.467	
	MIN, MAX	4410.200, 31587.800	12.833, 33.457	
	5%, 95%	4410.200, 31587.800	12.833, 33.457	

* P-value from wilcoxon rank-sum test.
* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

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TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 44 of ?)

Biomarker: Total Cotinine (ug/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	6	0.0051
	MEAN, GEO MEAN	3795.140, 3124.500	3.960, 3.674	
	SD, SE	1978.1800, 807.5880	1.6917, 0.6906	
	MEDIAN	3849.890	3.412	
	MIN, MAX	686.761, 6102.113	2.326, 6.218	
	5%, 95%	686.761, 6102.110	2.326, 6.218	
	BLQ Values Included	0	6	
Week 2	N	6	7	0.0034
	MEAN, GEO MEAN	3866.320, 3492.440	4.520, 4.212	
	SD, SE	1524.9400, 622.5540	1.6696, 0.6310	
	MEDIAN	4421.120	5.330	
	MIN, MAX	1214.933, 5109.091	2.016, 6.590	
	5%, 95%	1214.930, 5109.090	2.016, 6.590	
	BLQ Values Included	0	7	
Week 3	N	5	7	0.0058
	MEAN, GEO MEAN	3235.010, 2406.610	3.955, 3.789	
	SD, SE	2018.3200, 902.6200	1.2545, 0.4742	
	MEDIAN	3596.320	3.644	
	MIN, MAX	429.535, 5537.141	2.442, 5.640	
	5%, 95%	429.535, 5537.140	2.442, 5.640	
	BLQ Values Included	0	7	
Overall*	N	6	8	0.8983
	MEAN, GEO MEAN	3678.750, 3120.270	4.158, 3.954	
	SD, SE	1755.1600, 716.5410	1.3489, 0.4769	
	MEDIAN	4035.340	4.135	
	MIN, MAX	777.076, 5565.770	2.261, 5.895	
	5%, 95%	777.076, 5565.770	2.261, 5.895	

[a] P-value from wilcoxon rank-sum test.

* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

PM3003565268

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Free Cotinine (nmol/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	7	0.0034
	MEAN, GEO MEAN	8116.240, 7377.370	9.819, 7.923	
	SD, SE	3337.3900, 1362.4800	8.2449, 3.1163	
	MEDIAN	9239.020	7.062	
	MIN, MAX	3100.528, 11659.550	3.720, 27.797	
	5%, 95%	3100.530, 11659.600	3.720, 27.797	
	BLQ Values Included	0	6	
Week 2	N	6	8	0.0024
	MEAN, GEO MEAN	8657.220, 8169.360	7.179, 6.747	
	SD, SE	3203.9800, 1308.0200	2.4766, 0.8756	
	MEDIAN	8162.920	7.673	
	MIN, MAX	4574.775, 13936.670	3.224, 10.540	
	5%, 95%	4574.780, 13936.700	3.224, 10.540	
	BLQ Values Included	0	8	
Week 3	N	5	7	0.0058
	MEAN, GEO MEAN	7331.940, 6175.170	6.454, 6.143	
	SD, SE	4110.3900, 1838.2200	2.2246, 0.8408	
	MEDIAN	7564.870	5.828	
	MIN, MAX	2098.704, 12059.465	3.906, 9.920	
	5%, 95%	2098.700, 12059.500	3.906, 9.920	
	BLQ Values Included	0	7	
Overall[b]	N	6	8	1.0000
	MEAN, GEO MEAN	8075.000, 7437.280	7.852, 7.377	
	SD, SE	3105.3600, 1267.7600	2.8719, 1.0154	
	MEDIAN	9222.000	7.854	
	MIN, MAX	3258.000, 10736.400	3.617, 13.134	
	5%, 95%	3258.000, 10736.400	3.617, 13.134	

[a] P-value from wilcoxon rank-sum test.
[b] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminating all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 46 of ?)

Biomarker: Free Cotinine (ug/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	7	0.0034
	MEAN, GEO MEAN	1430.080, 1299.890	1.730, 1.396	
	SD, SE	588.0490, 240.0700	1.4529, 0.5492	
	MEDIAN	1627.920	1.244	
	MIN, MAX	546.313, 2054.413	0.655, 4.898	
	5%, 95%	546.313, 2054.410	0.655, 4.898	
	BLQ Values Included	0	6	
Week 2	N	6	8	0.0024
	MEAN, GEO MEAN	1525.400, 1439.440	1.265, 1.189	
	SD, SE	564.5420, 230.4730	0.4362, 0.1542	
	MEDIAN	1438.310	1.352	
	MIN, MAX	806.075, 2455.641	0.568, 1.857	
	5%, 95%	806.075, 2455.640	0.568, 1.857	
	BLQ Values Included	0	8	
Week 3	N	5	7	0.0058
	MEAN, GEO MEAN	1291.890, 1088.070	1.137, 1.082	
	SD, SE	724.2500, 323.8950	0.3920, 0.1482	
	MEDIAN	1332.930	1.027	
	MIN, MAX	369.792, 2124.878	0.688, 1.748	
	5%, 95%	369.792, 2124.880	0.688, 1.748	
	BLQ Values Included	0	7	
Overall[*]	N	6	8	1.0000
	MEAN, GEO MEAN	1422.820, 1310.450	1.383, 1.300	
	SD, SE	547.1660, 223.3790	0.5060, 0.1789	
	MEDIAN	1624.920	1.384	
	MIN, MAX	574.060, 1891.760	0.637, 2.314	
	5%, 95%	574.060, 1891.760	0.637, 2.314	

[a] P-value from wilcoxon rank-sum test.
[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

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TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 47 of ?)

Biomarker: Total 3-hydroxycotinine (nmol/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	5	0.0081
	MEAN, GEO MEAN	33202.000, 29446.300	54.413, 41.136	
	SD, SE	16444.0000, 6713.2200	46.9354, 20.9901	
	MEDIAN	32778.000	39.779	
	MIN, MAX	11447.120, 59281.220	14.880, 132.925	
	5%, 95%	11447.100, 59281.200	14.880, 132.925	
	BLQ Values Included	0	3	
Week 2	N	6	5	0.0081
	MEAN, GEO MEAN	34757.100, 29644.700	35.146, 29.986	
	SD, SE	23109.7000, 9434.5000	22.0406, 9.8569	
	MEDIAN	28940.000	35.650	
	MIN, MAX	13685.048, 78161.850	12.896, 70.392	
	5%, 95%	13685.000, 78161.900	12.896, 70.392	
	BLQ Values Included	0	4	
Week 3	N	5	5	0.0122
	MEAN, GEO MEAN	25853.900, 18736.600	27.836, 26.364	
	SD, SE	24683.3000, 11038.7000	9.8852, 4.4208	
	MEDIAN	16042.200	25.594	
	MIN, MAX	5906.262, 68488.280	15.624, 39.680	
	5%, 95%	5906.260, 68488.300	15.624, 39.680	
	BLQ Values Included	0	5	
Overall*	N	6	6	0.6889
	MEAN, GEO MEAN	32057.000, 27264.900	49.377, 38.249	
	SD, SE	20295.1000, 8285.4600	42.9212, 17.5225	
	MEDIAN	25716.000	37.165	
	MIN, MAX	10346.100, 68643.800	14.467, 132.925	
	5%, 95%	10346.100, 68643.800	14.467, 132.925	

[a] P-value from wilcoxon rank-sum test.
* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

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TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Total 3-hydroxycotinine (ug/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	5	0.0081
	MEAN, GEO MEAN	6381.420, 5659.580	10.458, 7.907	
	SD, SE	3160.5300, 1290.2800	9.0208, 4.0342	
	MEDIAN	6299.940	7.646	
	MIN, MAX	2200.136, 11393.850	2.860, 25.548	
	5%, 95%	2200.140, 11393.900	2.860, 25.548	
	BLQ Values Included	0	3	
Week 2	N	6	5	0.0081
	MEAN, GEO MEAN	6680.320, 5697.710	6.755, 5.764	
	SD, SE	4441.6900, 1813.3100	4.2360, 1.8944	
	MEDIAN	5562.270	6.852	
	MIN, MAX	2630.266, 15022.708	2.479, 13.529	
	5%, 95%	2630.270, 15022.700	2.479, 13.529	
	BLQ Values Included	0	4	
Week 3	N	5	5	0.0122
	MEAN, GEO MEAN	4969.130, 3601.180	5.350, 5.067	
	SD, SE	4744.1200, 2121.6400	1.8999, 0.8496	
	MEDIAN	3083.320	4.919	
	MIN, MAX	1135.184, 13163.447	3.003, 7.626	
	5%, 95%	1135.180, 13163.400	3.003, 7.626	
	BLQ Values Included	0	5	
Overall[*]	N	6	6	0.6889
	MEAN, GEO MEAN	6161.340, 5240.300	9.490, 7.352	
	SD, SE	3900.7100, 1592.4600	8.2493, 3.3678	
	MEDIAN	4942.610	7.144	
	MIN, MAX	1988.530, 13193.300	2.781, 25.548	
	5%, 95%	1988.530, 13193.300	2.781, 25.548	

[a] P-value from wilcoxon rank-sum test.
[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminating all BLQ values for the overall statistics.

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TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Free 3-hydroxycotinine (nmol/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	8	0.0024
	MEAN, GEO MEAN	27286.800, 24423.200	27.743, 20.815	
	SD, SE	13018.2000, 5314.6700	19.8790, 7.0283	
	MEDIAN	26551.000	23.760	
	MIN, MAX	9903.928, 47408.910	5.880, 56.550	
	5%, 95%	9903.930, 47408.900	5.880, 56.550	
	BLQ Values Included	0	0	
Week 2	N	6	7	0.0034
	MEAN, GEO MEAN	28780.800, 23570.900	26.811, 22.854	
	SD, SE	20864.7000, 8517.9700	15.6177, 5.9029	
	MEDIAN	24714.300	23.120	
	MIN, MAX	9062.489, 67799.170	10.712, 47.850	
	5%, 95%	9062.490, 67799.200	10.712, 47.850	
	BLQ Values Included	0	0	
Week 3	N	5	5	0.0122
	MEAN, GEO MEAN	20684.400, 15880.900	22.055, 19.578	
	SD, SE	17932.5000, 8019.6800	11.5204, 5.1521	
	MEDIAN	13910.500	18.228	
	MIN, MAX	5533.662, 51582.300	9.450, 36.375	
	5%, 95%	5533.660, 51582.300	9.450, 36.375	
	BLQ Values Included	0	0	
Overall[b]	N	6	8	0.8983
	MEAN, GEO MEAN	26279.500, 22293.300	23.547, 19.730	
	SD, SE	16487.3000, 6730.9100	13.4248, 4.7464	
	MEDIAN	21636.100	24.725	
	MIN, MAX	8166.690, 55596.800	8.681, 38.675	
	5%, 95%	8166.690, 55596.800	8.681, 38.675	

[a] P-value from wilcoxon rank-sum test.

[b] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

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TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 50 of ?)

Biomarker: Free 3-hydroxycotinine (ug/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value*
Week 1	N	6	8	0.0024
	MEAN, GEO MEAN	5244.520, 4694.150	5.332, 4.000	
	SD, SE	2502.1100, 1021.4800	3.8207, 1.3508	
	MEDIAN	5103.110	4.567	
	MIN, MAX	1903.535, 9111.993	1.130, 10.869	
	5%, 95%	1903.540, 9111.990	1.130, 10.869	
	BLQ Values Included	0	0	
Week 2	N	6	7	0.0034
	MEAN, GEO MEAN	5531.660, 4530.330	5.153, 4.393	
	SD, SE	4010.1900, 1637.1500	3.0018, 1.1346	
	MEDIAN	4750.100	4.444	
	MIN, MAX	1741.810, 13031.000	2.059, 9.197	
	5%, 95%	1741.810, 13031.000	2.059, 9.197	
	BLQ Values Included	0	0	
Week 3	N	5	5	0.0122
	MEAN, GEO MEAN	3975.540, 3052.300	4.239, 3.763	
	SD, SE	3446.6400, 1541.3800	2.2143, 0.9903	
	MEDIAN	2673.600	3.503	
	MIN, MAX	1063.570, 9914.118	1.816, 6.991	
	5%, 95%	1063.570, 9914.120	1.816, 6.991	
	BLQ Values Included	0	0	
Overall*	N	6	8	0.8983
	MEAN, GEO MEAN	5050.920, 4284.770	4.526, 3.792	
	SD, SE	3168.8600, 1293.6800	2.5804, 0.9123	
	MEDIAN	4158.460	4.752	
	MIN, MAX	1569.640, 10685.700	1.668, 7.433	
	5%, 95%	1569.640, 10685.700	1.668, 7.433	

* P-value from wilcoxon rank-sum test.
* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Total Nicotine (nmol/mg Creatin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	3	0.0282
	MEAN, GEO MEAN	9.381, 7.928	0.024, 0.022	
	SD, SE	5.4803, 2.2373	0.0129, 0.0074	
	MEDIAN	8.947	0.019	
	MIN, MAX	2.896, 17.795	0.015, 0.039	
	5%, 95%	2.896, 17.795	0.015, 0.039	
	BLQ Values Included	0	3	
Week 2	N	6	5	0.0081
	MEAN, GEO MEAN	9.755, 8.845	0.031, 0.029	
	SD, SE	4.6862, 1.9131	0.0118, 0.0053	
	MEDIAN	9.474	0.027	
	MIN, MAX	4.131, 17.906	0.016, 0.044	
	5%, 95%	4.131, 17.906	0.016, 0.044	
	BLQ Values Included	0	4	
Week 3	N	5	7	0.0057
	MEAN, GEO MEAN	8.488, 7.874	0.021, 0.020	
	SD, SE	3.6972, 1.6534	0.0064, 0.0024	
	MEDIAN	6.628	0.024	
	MIN, MAX	4.861, 13.377	0.013, 0.029	
	5%, 95%	4.861, 13.377	0.013, 0.029	
	BLQ Values Included	0	7	
Overall[b]	N	6	7	0.7210
	MEAN, GEO MEAN	9.367, 8.525	0.025, 0.024	
	SD, SE	4.2474, 1.7340	0.0072, 0.0027	
	MEDIAN	9.697	0.025	
	MIN, MAX	4.390, 15.711	0.015, 0.034	
	5%, 95%	4.390, 15.711	0.015, 0.034	

[a] P-value from wilcoxon rank-sum test.
[b] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminating all BLQ values for the overall statistics.

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TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Total Nicotine (ug/mg Creatinin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	3	0.0282
	MEAN, GEO MEAN	1.522, 1.286	0.004, 0.003	
	SD, SE	0.8887, 0.3628	0.0021, 0.0012	
	MEDIAN	1.451	0.003	
	MIN, MAX	0.470, 2.886	0.002, 0.006	
	5%, 95%	0.470, 2.886	0.002, 0.006	
	BLQ Values Included	0	3	
Week 2	N	6	5	0.0078
	MEAN, GEO MEAN	1.582, 1.434	0.005, 0.005	
	SD, SE	0.7601, 0.3103	0.0019, 0.0008	
	MEDIAN	1.537	0.004	
	MIN, MAX	0.670, 2.904	0.003, 0.007	
	5%, 95%	0.670, 2.904	0.003, 0.007	
	BLQ Values Included	0	4	
Week 3	N	5	7	0.0053
	MEAN, GEO MEAN	1.377, 1.277	0.003, 0.003	
	SD, SE	0.5999, 0.2683	0.0011, 0.0004	
	MEDIAN	1.075	0.004	
	MIN, MAX	0.788, 2.170	0.002, 0.005	
	5%, 95%	0.788, 2.170	0.002, 0.005	
	BLQ Values Included	0	7	
Overall[b]	N	6	7	0.6151
	MEAN, GEO MEAN	1.519, 1.383	0.004, 0.004	
	SD, SE	0.6888, 0.2812	0.0013, 0.0005	
	MEDIAN	1.573	0.004	
	MIN, MAX	0.712, 2.548	0.002, 0.006	
	5%, 95%	0.712, 2.548	0.002, 0.006	

[a] P-value from wilcoxon rank-sum test.
[b] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 53 of ?)

Biomarker: Free Nicotine (nmol/mg Creatin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	3	0.0275
	MEAN, GEO MEAN	6.631, 5.419	0.005, 0.005	
	SD, SE	4.3231, 1.7649	0.0006, 0.0003	
	MEDIAN	5.003	0.005	
	MIN, MAX	1.717, 12.677	0.004, 0.005	
	5%, 95%	1.717, 12.677	0.004, 0.005	
	BLQ Values Included	0	3	
Week 2	N	6	2	0.0668
	MEAN, GEO MEAN	6.414, 5.576	0.008, 0.006	
	SD, SE	3.4787, 1.4202	0.0064, 0.0045	
	MEDIAN	6.483	0.008	
	MIN, MAX	2.159, 12.067	0.003, 0.012	
	5%, 95%	2.159, 12.067	0.003, 0.012	
	BLQ Values Included	0	2	
Week 3	N	5	4	0.0200
	MEAN, GEO MEAN	5.449, 5.159	0.006, 0.005	
	SD, SE	2.0593, 0.9210	0.0024, 0.0012	
	MEDIAN	4.773	0.006	
	MIN, MAX	3.605, 8.447	0.003, 0.008	
	5%, 95%	3.605, 8.447	0.003, 0.008	
	BLQ Values Included	0	4	
Overall*	N	6	6	0.9361
	MEAN, GEO MEAN	6.376, 5.742	0.006, 0.006	
	SD, SE	3.0639, 1.2508	0.0032, 0.0013	
	MEDIAN	5.903	0.005	
	MIN, MAX	2.883, 10.457	0.004, 0.012	
	5%, 95%	2.883, 10.457	0.004, 0.012	

[a] P-value from wilcoxon rank-sum test.

* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 54 of ?)

Biomarker: Free Nicotine (ug/mg Creatinin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	3	0.0256
	MEAN, GEO MEAN	1.076, 0.879	0.001, 0.001	
	SD, SE	0.7014, 0.2863	0.0000, 0.0000	
	MEDIAN	0.812	0.001	
	MIN, MAX	0.278, 2.056	0.001, 0.001	
	5%, 95%	0.278, 2.056	0.001, 0.001	
	BLQ Values Included	0	3	
Week 2	N	6	2	0.0668
	MEAN, GEO MEAN	1.040, 0.904	0.001, NA	
	SD, SE	0.5642, 0.2303	0.0014, 0.0010	
	MEDIAN	1.052	0.001	
	MIN, MAX	0.350, 1.957	0.000, 0.002	
	5%, 95%	0.350, 1.957	0.000, 0.002	
	BLQ Values Included	0	2	
Week 3	N	5	4	0.0179
	MEAN, GEO MEAN	0.884, 0.837	0.001, NA	
	SD, SE	0.3339, 0.1493	0.0005, 0.0003	
	MEDIAN	0.774	0.001	
	MIN, MAX	0.585, 1.370	0.000, 0.001	
	5%, 95%	0.585, 1.370	0.000, 0.001	
	BLQ Values Included	0	4	
Overall[b]	N	6	6	0.7444
	MEAN, GEO MEAN	1.034, 0.931	0.001, NA	
	SD, SE	0.4971, 0.2029	0.0006, 0.0003	
	MEDIAN	0.958	0.001	
	MIN, MAX	0.467, 1.696	0.000, 0.002	
	5%, 95%	0.467, 1.696	0.000, 0.002	

[a] P-value from wilcoxon rank-sum test.
[b] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Total Cotinine (nmol/mg Creatin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value*
Week 1	N	6	6	0.0050
	MEAN, GEO MEAN	15.933, 12.847	0.016, 0.014	
	SD, SE	9.1696, 3.7435	0.0098, 0.0040	
	MEDIAN	17.813	0.015	
	MIN, MAX	3.209, 24.408	0.006, 0.034	
	5%, 95%	3.209, 24.408	0.006, 0.034	
	BLQ Values Included	0	6	
Week 2	N	6	7	0.0034
	MEAN, GEO MEAN	17.374, 15.619	0.018, 0.016	
	SD, SE	8.1799, 3.3394	0.0107, 0.0041	
	MEDIAN	16.762	0.014	
	MIN, MAX	7.201, 27.449	0.007, 0.038	
	5%, 95%	7.201, 27.449	0.007, 0.038	
	BLQ Values Included	0	7	
Week 3	N	5	7	0.0056
	MEAN, GEO MEAN	16.373, 13.344	0.018, 0.017	
	SD, SE	9.6689, 4.3241	0.0051, 0.0019	
	MEDIAN	19.648	0.021	
	MIN, MAX	4.351, 25.375	0.011, 0.023	
	5%, 95%	4.351, 25.375	0.011, 0.023	
	BLQ Values Included	0	7	
Overall*	N	6	8	0.5224
	MEAN, GEO MEAN	16.432, 14.120	0.019, 0.018	
	SD, SE	8.4653, 3.4559	0.0077, 0.0027	
	MEDIAN	17.962	0.018	
	MIN, MAX	4.920, 25.193	0.009, 0.030	
	5%, 95%	4.920, 25.193	0.009, 0.030	

* P-value from wilcoxon rank-sum test.
* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

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TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 56 of ?)

Biomarker: Total Cotinine (ug/mg Creatinin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	6	0.0049
	MEAN, GEO MEAN	2.807, 2.263	0.003, 0.002	
	SD, SE	1.6159, 0.6597	0.0017, 0.0007	
	MEDIAN	3.139	0.003	
	MIN, MAX	0.565, 4.301	0.001, 0.006	
	5%, 95%	0.565, 4.301	0.001, 0.006	
	BLQ Values Included	0	6	
Week 2	N	6	7	0.0032
	MEAN, GEO MEAN	3.061, 2.752	0.003, 0.003	
	SD, SE	1.4415, 0.5885	0.0020, 0.0008	
	MEDIAN	2.954	0.002	
	MIN, MAX	1.269, 4.837	0.001, 0.007	
	5%, 95%	1.269, 4.837	0.001, 0.007	
	BLQ Values Included	0	7	
Week 3	N	5	7	0.0046
	MEAN, GEO MEAN	2.885, 2.352	0.003, 0.003	
	SD, SE	1.7035, 0.7618	0.0011, 0.0004	
	MEDIAN	3.462	0.004	
	MIN, MAX	0.767, 4.471	0.002, 0.004	
	5%, 95%	0.767, 4.471	0.002, 0.004	
	BLQ Values Included	0	7	
Overall[*]	N	6	8	0.4433
	MEAN, GEO MEAN	2.896, 2.488	0.003, 0.003	
	SD, SE	1.4915, 0.6089	0.0017, 0.0006	
	MEDIAN	3.165	0.003	
	MIN, MAX	0.867, 4.439	0.001, 0.006	
	5%, 95%	0.867, 4.439	0.001, 0.006	

[a] P-value from wilcoxon rank-sum test.

[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Free Cotinine (nmol/mg Creatin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	7	0.0032
	MEAN, GEO MEAN	6.206, 5.345	0.006, 0.006	
	SD, SE	3.4820, 1.4215	0.0034, 0.0013	
	MEDIAN	6.151	0.005	
	MIN, MAX	2.553, 11.433	0.003, 0.012	
	5%, 95%	2.553, 11.433	0.003, 0.012	
	BLQ Values Included	0	6	
Week 2	N	6	8	0.0024
	MEAN, GEO MEAN	7.082, 6.437	0.005, 0.005	
	SD, SE	3.7617, 1.5357	0.0028, 0.0010	
	MEDIAN	5.586	0.005	
	MIN, MAX	4.231, 14.182	0.002, 0.011	
	5%, 95%	4.231, 14.182	0.002, 0.011	
	BLQ Values Included	0	8	
Week 3	N	5	7	0.0049
	MEAN, GEO MEAN	6.752, 6.033	0.005, 0.005	
	SD, SE	3.2634, 1.4594	0.0014, 0.0005	
	MEDIAN	7.282	0.006	
	MIN, MAX	3.016, 9.978	0.003, 0.007	
	5%, 95%	3.016, 9.978	0.003, 0.007	
	BLQ Values Included	0	7	
Overall[*]	N	6	8	0.7012
	MEAN, GEO MEAN	6.575, 5.962	0.006, 0.006	
	SD, SE	3.0766, 1.2560	0.0018, 0.0006	
	MEDIAN	6.547	0.005	
	MIN, MAX	3.296, 10.966	0.004, 0.009	
	5%, 95%	3.296, 10.966	0.004, 0.009	

[a] P-value from wilcoxon rank-sum test.

[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Free Cotinine (ug/mg Creatinin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	7	0.0026
	MEAN, GEO MEAN	1.093, 0.942	0.001, 0.001	
	SD, SE	0.6133, 0.2504	0.0005, 0.0002	
	MEDIAN	1.084	0.001	
	MIN, MAX	0.450, 2.014	0.001, 0.002	
	5%, 95%	0.450, 2.014	0.001, 0.002	
	BLQ Values Included	0	6	
Week 2	N	6	8	0.0016
	MEAN, GEO MEAN	1.248, 1.134	0.001, NA	
	SD, SE	0.6628, 0.2706	0.0005, 0.0002	
	MEDIAN	0.984	0.001	
	MIN, MAX	0.746, 2.499	0.000, 0.002	
	5%, 95%	0.746, 2.499	0.000, 0.002	
	BLQ Values Included	0	8	
Week 3	N	5	7	0.0021
	MEAN, GEO MEAN	1.190, 1.063	0.001, 0.001	
	SD, SE	0.5751, 0.2572	0.0000, 0.0000	
	MEDIAN	1.283	0.001	
	MIN, MAX	0.531, 1.758	0.001, 0.001	
	5%, 95%	0.531, 1.758	0.001, 0.001	
	BLQ Values Included	0	7	
Overall[*]	N	6	8	0.5947
	MEAN, GEO MEAN	1.159, 1.051	0.001, 0.001	
	SD, SE	0.5419, 0.2212	0.0004, 0.0001	
	MEDIAN	1.154	0.001	
	MIN, MAX	0.581, 1.932	0.001, 0.002	
	5%, 95%	0.581, 1.932	0.001, 0.002	

[a] P-value from wilcoxon rank-sum test.

[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2

STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Total 3-hydroxycotinine (nmol/mg Creatin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	5	0.0080
	MEAN, GEO MEAN	26.386, 21.334	0.040, 0.032	
	SD, SE	21.9570, 8.9639	0.0326, 0.0146	
	MEDIAN	20.188	0.025	
	MIN, MAX	9.426, 70.006	0.019, 0.096	
	5%, 95%	9.426, 70.006	0.019, 0.096	
	BLQ Values Included	0	3	
Week 2	N	6	5	0.0075
	MEAN, GEO MEAN	28.860, 23.360	0.021, 0.020	
	SD, SE	25.1708, 10.2759	0.0073, 0.0032	
	MEDIAN	19.587	0.026	
	MIN, MAX	14.293, 79.538	0.011, 0.026	
	5%, 95%	14.293, 79.538	0.011, 0.026	
	BLQ Values Included	0	4	
Week 3	N	5	5	0.0122
	MEAN, GEO MEAN	24.228, 18.306	0.021, 0.020	
	SD, SE	23.6263, 10.5660	0.0070, 0.0031	
	MEDIAN	15.321	0.023	
	MIN, MAX	9.955, 65.930	0.012, 0.028	
	5%, 95%	9.955, 65.930	0.012, 0.028	
	BLQ Values Included	0	5	
Overall*	N	6	6	0.6889
	MEAN, GEO MEAN	26.496, 21.465	0.035, 0.028	
	SD, SE	22.6461, 9.2452	0.0306, 0.0125	
	MEDIAN	18.599	0.024	
	MIN, MAX	11.420, 71.825	0.014, 0.096	
	5%, 95%	11.420, 71.825	0.014, 0.096	

[a] P-value from wilcoxon rank-sum test.

* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 60 of ?)

Biomarker: Total 3-hydroxycotinine (ug/mg Creatinin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value*
Week 1	N	6	5	0.0080
	MEAN, GEO MEAN	5.071, 4.100	0.008, 0.006	
	SD, SE	4.2201, 1.7228	0.0059, 0.0027	
	MEDIAN	3.880	0.005	
	MIN, MAX	1.812, 13.455	0.004, 0.018	
	5%, 95%	1.812, 13.455	0.004, 0.018	
	BLQ Values Included	0	3	
Week 2	N	6	5	0.0075
	MEAN, GEO MEAN	5.547, 4.490	0.004, 0.004	
	SD, SE	4.8377, 1.9750	0.0014, 0.0006	
	MEDIAN	3.765	0.005	
	MIN, MAX	2.747, 15.287	0.002, 0.005	
	5%, 95%	2.747, 15.287	0.002, 0.005	
	BLQ Values Included	0	4	
Week 3	N	5	5	0.0119
	MEAN, GEO MEAN	4.657, 3.518	0.004, 0.004	
	SD, SE	4.5411, 2.0309	0.0013, 0.0006	
	MEDIAN	2.945	0.004	
	MIN, MAX	1.913, 12.672	0.002, 0.005	
	5%, 95%	1.913, 12.672	0.002, 0.005	
	BLQ Values Included	0	5	
Overall*	N	6	6	0.6889
	MEAN, GEO MEAN	5.093, 4.126	0.007, 0.005	
	SD, SE	4.3527, 1.7770	0.0056, 0.0023	
	MEDIAN	3.575	0.005	
	MIN, MAX	2.195, 13.805	0.003, 0.018	
	5%, 95%	2.195, 13.805	0.003, 0.018	

[a] P-value from wilcoxon rank-sum test.
* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 61 of 7)

Biomarker: Free 3-hydroxycotinine (nmol/mg Creatin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	8	0.0024
	MEAN, GEO MEAN	21.591, 17.695	0.018, 0.014	
	SD, SE	17.3519, 7.0839	0.0129, 0.0046	
	MEDIAN	16.395	0.014	
	MIN, MAX	8.155, 55.986	0.005, 0.041	
	5%, 95%	8.155, 55.986	0.005, 0.041	
	BLQ Values Included	0	0	
Week 2	N	6	7	0.0034
	MEAN, GEO MEAN	23.863, 18.574	0.016, 0.015	
	SD, SE	22.4716, 9.1740	0.0066, 0.0025	
	MEDIAN	15.847	0.017	
	MIN, MAX	9.465, 68.993	0.008, 0.024	
	5%, 95%	9.465, 68.993	0.008, 0.024	
	BLQ Values Included	0	0	
Week 3	N	5	5	0.0119
	MEAN, GEO MEAN	19.402, 15.516	0.018, 0.015	
	SD, SE	17.0839, 7.6401	0.0108, 0.0048	
	MEDIAN	12.774	0.016	
	MIN, MAX	9.346, 49.656	0.008, 0.032	
	5%, 95%	9.346, 49.656	0.008, 0.032	
	BLQ Values Included	0	0	
Overall[*]	N	6	8	0.6093
	MEAN, GEO MEAN	21.660, 17.595	0.016, 0.013	
	SD, SE	18.2969, 7.4697	0.0094, 0.0033	
	MEDIAN	15.202	0.014	
	MIN, MAX	9.166, 58.212	0.005, 0.032	
	5%, 95%	9.166, 58.212	0.005, 0.032	

[a] P-value from wilcoxon rank-sum test.

[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

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TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Free 3-hydroxycotinine (ug/mg Creatinin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	8	0.0023
	MEAN, GEO MEAN	4.150, 3.401	0.003, 0.003	
	SD, SE	3.3353, 1.3616	0.0025, 0.0009	
	MEDIAN	3.151	0.003	
	MIN, MAX	1.567, 10.761	0.001, 0.008	
	5%, 95%	1.567, 10.761	0.001, 0.008	
	BLQ Values Included	0	0	
Week 2	N	6	7	0.0032
	MEAN, GEO MEAN	4.586, 3.570	0.003, 0.003	
	SD, SE	4.3189, 1.7632	0.0012, 0.0005	
	MEDIAN	3.046	0.003	
	MIN, MAX	1.819, 13.260	0.002, 0.005	
	5%, 95%	1.819, 13.260	0.002, 0.005	
	BLQ Values Included	0	0	
Week 3	N	5	5	0.0119
	MEAN, GEO MEAN	3.729, 2.982	0.004, 0.003	
	SD, SE	3.2837, 1.4685	0.0018, 0.0008	
	MEDIAN	2.455	0.003	
	MIN, MAX	1.796, 9.544	0.002, 0.006	
	5%, 95%	1.796, 9.544	0.002, 0.006	
	BLQ Values Included	0	0	
Overall[*]	N	6	8	0.6093
	MEAN, GEO MEAN	4.163, 3.382	0.003, 0.003	
	SD, SE	3.5166, 1.4356	0.0016, 0.0006	
	MEDIAN	2.922	0.003	
	MIN, MAX	1.761, 11.188	0.001, 0.006	
	5%, 95%	1.761, 11.188	0.001, 0.006	

[a] P-value from wilcoxon rank-sum test.

[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

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TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Nicotine Gluruonide (nmol/L)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	2	0.0651
	MEAN, GEO MEAN	2601.8, 1698.4	9.3, 9.3	
	SD, SE	2910.33, 1188.14	0.00, 0.00	
	MEDIAN	1692.2	9.3	
	MIN, MAX	403.4, 8373.6	9.3, 9.3	
	5%, 95%	403.4, 8373.6	9.3, 9.3	
	BLQ Values Included	0	2	
Week 2	N	6	2	0.0668
	MEAN, GEO MEAN	2877.9, 2365.5	27.7, 20.7	
	SD, SE	2171.62, 886.56	25.95, 18.35	
	MEDIAN	2040.6	27.7	
	MIN, MAX	1145.6, 6965.8	9.3, 46.0	
	5%, 95%	1145.6, 6965.8	9.3, 46.0	
	BLQ Values Included	0	2	
Week 3	N	5	4	0.0151
	MEAN, GEO MEAN	1850.7, 1460.3	9.3, 9.3	
	SD, SE	1312.92, 587.16	0.00, 0.00	
	MEDIAN	1299.6	9.3	
	MIN, MAX	432.7, 3839.2	9.3, 9.3	
	5%, 95%	432.7, 3839.2	9.3, 9.3	
	BLQ Values Included	0	4	
Overall*	N	6	6	1.0000
	MEAN, GEO MEAN	2495.4, 1962.5	11.3, 10.7	
	SD, SE	2051.46, 837.50	4.98, 2.03	
	MEDIAN	1908.5	9.3	
	MIN, MAX	797.4, 6392.9	9.3, 21.5	
	5%, 95%	797.4, 6392.9	9.3, 21.5	

[a] P-value from wilcoxon rank-sum test.
* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

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TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Nicotine Gluruonide (nmol/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	2	0.0668
	MEAN, GEO MEAN	3721.250, 2858.730	19.530, 17.646	
	SD, SE	2425.2300, 990.0960	11.8370, 8.3700	
	MEDIAN	3264.810	19.530	
	MIN, MAX	548.624, 7285.032	11.160, 27.900	
	5%, 95%	548.624, 7285.030	11.160, 27.900	
	BLQ Values Included	0	2	
Week 2	N	6	2	0.0668
	MEAN, GEO MEAN	4160.740, 3985.670	39.730, 38.893	
	SD, SE	1399.0600, 571.1660	11.4693, 8.1100	
	MEDIAN	3587.760	39.730	
	MIN, MAX	2816.360, 6478.194	31.620, 47.840	
	5%, 95%	2816.360, 6478.190	31.620, 47.840	
	BLQ Values Included	0	2	
Week 3	N	5	4	0.0200
	MEAN, GEO MEAN	3377.800, 2594.130	19.353, 18.296	
	SD, SE	2201.0600, 984.3430	7.6038, 3.8019	
	MEDIAN	3139.430	17.968	
	MIN, MAX	597.126, 5950.760	11.718, 29.760	
	5%, 95%	597.126, 5950.760	11.718, 29.760	
	BLQ Values Included	0	4	
Overall*	N	6	6	0.6889
	MEAN, GEO MEAN	3758.070, 3321.290	24.798, 24.160	
	SD, SE	1913.2300, 781.0710	5.9768, 2.4400	
	MEDIAN	3384.820	25.737	
	MIN, MAX	1320.700, 6571.330	16.740, 31.620	
	5%, 95%	1320.700, 6571.330	16.740, 31.620	

[a] P-value from wilcoxon rank-sum test.

* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 65 of ?)

Biomarker: Nicotine Gluruonide (nmol/mg Creatin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	2	0.0668
	MEAN, GEO MEAN	2.750, 2.071	0.013, 0.012	
	SD, SE	1.8455, 0.7534	0.0021, 0.0015	
	MEDIAN	2.837	0.013	
	MIN, MAX	0.452, 5.118	0.011, 0.014	
	5%, 95%	0.452, 5.118	0.011, 0.014	
	BLQ Values Included	0	2	
Week 2	N	6	2	0.0668
	MEAN, GEO MEAN	3.341, 3.141	0.036, 0.036	
	SD, SE	1.3634, 0.5566	0.0042, 0.0030	
	MEDIAN	3.126	0.036	
	MIN, MAX	1.972, 5.839	0.033, 0.039	
	5%, 95%	1.972, 5.839	0.033, 0.039	
	BLQ Values Included	0	2	
Week 3	N	5	4	0.0200
	MEAN, GEO MEAN	3.039, 2.535	0.015, 0.014	
	SD, SE	1.8280, 0.8175	0.0059, 0.0029	
	MEDIAN	3.022	0.015	
	MIN, MAX	1.066, 4.929	0.009, 0.021	
	5%, 95%	1.066, 4.929	0.009, 0.021	
	BLQ Values Included	0	4	
Overall[*]	N	6	6	0.4704
	MEAN, GEO MEAN	2.992, 2.662	0.019, 0.018	
	SD, SE	1.5386, 0.6281	0.0081, 0.0033	
	MEDIAN	2.800	0.020	
	MIN, MAX	1.486, 5.254	0.011, 0.033	
	5%, 95%	1.486, 5.254	0.011, 0.033	

[a] P-value from wilcoxon rank-sum test.
[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Cotinine Gluruonide (nmol/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	5	0.0081
	MEAN, GEO MEAN	13422.600, 9233.170	17.266, 16.074	
	SD, SE	8208.4800, 3351.1000	7.0648, 3.1595	
	MEDIAN	12610.500	16.116	
	MIN, MAX	797.096, 24184.869	9.480, 25.343	
	5%, 95%	797.096, 24184.900	9.480, 25.343	
	BLQ Values Included	0	5	
Week 2	N	6	7	0.0034
	MEAN, GEO MEAN	13285.500, 10994.900	18.424, 17.167	
	SD, SE	6499.2400, 2653.3000	6.8053, 2.5722	
	MEDIAN	14471.800	21.725	
	MIN, MAX	2320.420, 19683.264	8.216, 26.860	
	5%, 95%	2320.420, 19683.300	8.216, 26.860	
	BLQ Values Included	0	7	
Week 3	N	5	6	0.0081
	MEAN, GEO MEAN	11027.900, 6237.610	14.976, 14.454	
	SD, SE	7390.1600, 3304.9800	4.5130, 1.8424	
	MEDIAN	12845.600	14.536	
	MIN, MAX	339.066, 19365.855	9.954, 22.910	
	5%, 95%	339.066, 19365.900	9.954, 22.910	
	BLQ Values Included	0	6	
Overall[*]	N	6	8	0.8983
	MEAN, GEO MEAN	12803.200, 9518.730	17.115, 16.315	
	SD, SE	7083.9900, 2892.0300	5.3727, 1.8995	
	MEDIAN	13680.100	16.980	
	MIN, MAX	1152.190, 21078.000	9.217, 24.028	
	5%, 95%	1152.190, 21078.000	9.217, 24.028	

[a] P-value from wilcoxon rank-sum test.

[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

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TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Cotinine Gluruonide (nmol/mg Creatin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	5	0.0080
	MEAN, GEO MEAN	9.727, 6.689	0.013, 0.012	
	SD, SE	6.1408, 2.5070	0.0070, 0.0031	
	MEDIAN	10.503	0.012	
	MIN, MAX	0.656, 16.992	0.007, 0.025	
	5%, 95%	0.656, 16.992	0.007, 0.025	
	BLQ Values Included	0	5	
Week 2	N	6	7	0.0034
	MEAN, GEO MEAN	10.292, 8.664	0.013, 0.011	
	SD, SE	5.4364, 2.2194	0.0079, 0.0030	
	MEDIAN	11.178	0.010	
	MIN, MAX	2.423, 17.741	0.005, 0.028	
	5%, 95%	2.423, 17.741	0.005, 0.028	
	BLQ Values Included	0	7	
Week 3	N	5	6	0.0078
	MEAN, GEO MEAN	9.621, 6.094	0.013, 0.013	
	SD, SE	6.5461, 2.9275	0.0037, 0.0015	
	MEDIAN	12.366	0.015	
	MIN, MAX	0.605, 15.637	0.008, 0.016	
	5%, 95%	0.605, 15.637	0.008, 0.016	
	BLQ Values Included	0	6	
Overall[b]	N	6	8	0.5229
	MEAN, GEO MEAN	9.857, 7.519	0.014, 0.013	
	SD, SE	5.7580, 2.3507	0.0055, 0.0019	
	MEDIAN	11.042	0.013	
	MIN, MAX	1.228, 16.790	0.007, 0.022	
	5%, 95%	1.228, 16.790	0.007, 0.022	

[a] P-value from wilcoxon rank-sum test.
[b] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

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TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Cotinine Glucuonide (nmol/L)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	5	0.0055
	MEAN, GEO MEAN	9217.7, 5485.6	7.9, 7.9	
	SD, SE	9594.01, 3916.74	0.00, 0.00	
	MEDIAN	7145.9	7.9	
	MIN, MAX	586.1, 27798.7	7.9, 7.9	
	5%, 95%	586.1, 27798.7	7.9, 7.9	
	BLQ Values Included	0	5	
Week 2	N	6	7	0.0015
	MEAN, GEO MEAN	9311.5, 6525.5	7.9, 7.9	
	SD, SE	7584.50, 3096.36	0.00, 0.00	
	MEDIAN	6886.9	7.9	
	MIN, MAX	1282.0, 21164.8	7.9, 7.9	
	5%, 95%	1282.0, 21164.8	7.9, 7.9	
	BLQ Values Included	0	7	
Week 3	N	5	6	0.0039
	MEAN, GEO MEAN	5950.2, 3511.4	7.9, 7.9	
	SD, SE	4438.14, 1984.80	0.00, 0.00	
	MEDIAN	5243.1	7.9	
	MIN, MAX	245.7, 12494.1	7.9, 7.9	
	5%, 95%	245.7, 12494.1	7.9, 7.9	
	BLQ Values Included	0	6	
Overall[*]	N	6	8	0.8875
	MEAN, GEO MEAN	8503.8, 5651.7	7.9, 7.9	
	SD, SE	7002.82, 2858.89	0.00, 0.00	
	MEDIAN	6639.0	7.9	
	MIN, MAX	704.6, 20485.9	7.9, 7.9	
	5%, 95%	704.6, 20485.9	7.9, 7.9	

[a] P-value from wilcoxon rank-sum test.
[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: 3-hydroxycotinine glucuronide (nmol/L)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	2	0.0668
	MEAN, GEO MEAN	3229.3, 2944.6	18.1, 17.2	
	SD, SE	1244.01, 507.87	7.71, 5.45	
	MEDIAN	3884.0	18.1	
	MIN, MAX	1134.7, 4122.8	12.6, 23.5	
	5%, 95%	1134.7, 4122.8	12.6, 23.5	
	BLQ Values Included	0	0	
Week 2	N	6	1	NA
	MEAN, GEO MEAN	3761.5, 3336.9	14.3, 14.3	
	SD, SE	2090.68, 853.52	NA, NA	
	MEDIAN	2948.4	14.3	
	MIN, MAX	1937.9, 7148.3	14.3, 14.3	
	5%, 95%	1937.9, 7148.3	NA, NA	
	BLQ Values Included	0	0	
Week 3	N	5		NA
	MEAN, GEO MEAN	2442.3, 1338.9		
	SD, SE	2744.27, 1227.28		
	MEDIAN	1240.5		
	MIN, MAX	270.0, 6900.4		
	5%, 95%	270.0, 6900.4		
	BLQ Values Included	0		
Overall*	N	6	2	1.0000
	MEAN, GEO MEAN	3314.7, 2908.5	18.5, 17.8	
	SD, SE	1730.35, 706.41	7.07, 5.00	
	MEDIAN	3199.8	18.5	
	MIN, MAX	1319.5, 5509.1	13.5, 23.5	
	5%, 95%	1319.5, 5509.1	13.5, 23.5	

[a] P-value from wilcoxon rank-sum test.
* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: 3-hydroxycotinine glucuronide (nmol/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	2	0.0668
	MEAN, GEO MEAN	5915.230, 4956.210	46.567, 35.776	
	SD, SE	3528.2700, 1440.4100	42.1556, 29.8085	
	MEDIAN	5879.110	46.567	
	MIN, MAX	1543.192, 11872.310	16.758, 76.375	
	5%, 95%	1543.190, 11872.300	16.758, 76.375	
	BLQ Values Included	0	0	
Week 2	N	6	1	NA
	MEAN, GEO MEAN	5976.360, 5622.440	24.024, 24.024	
	SD, SE	2460.4200, 1004.4600	NA, NA	
	MEDIAN	4840.690	24.024	
	MIN, MAX	3837.042, 10362.680	24.024, 24.024	
	5%, 95%	3837.040, 10362.700	NA, NA	
	BLQ Values Included	0	0	
Week 3	N	5		NA
	MEAN, GEO MEAN	5169.570, 2378.380		
	SD, SE	6804.7400, 3043.1700		
	MEDIAN	2667.080		
	MIN, MAX	372.600, 16905.980		
	5%, 95%	372.600, 16906.000		
	BLQ Values Included	0		
Overall[b]	N	6	2	0.8852
	MEAN, GEO MEAN	5777.450, 4931.670	48.383, 39.463	
	SD, SE	3856.0300, 1574.2200	39.5867, 27.9920	
	MEDIAN	4375.770	48.383	
	MIN, MAX	2179.450, 13047.000	20.391, 76.375	
	5%, 95%	2179.450, 13047.000	20.391, 76.375	

[a] P-value from wilcoxon rank-sum test.
[b] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminating all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: 3-hydroxycotinine glucuronide (nmol/mg Creatin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	2	0.0668
	MEAN, GEO MEAN	4.796, 3.591	0.031, 0.020	
	SD, SE	4.6266, 1.8888	0.0339, 0.0240	
	MEDIAN	3.582	0.031	
	MIN, MAX	1.271, 14.020	0.007, 0.055	
	5%, 95%	1.271, 14.020	0.007, 0.055	
	BLQ Values Included	0	0	
Week 2	N	6	1	NA
	MEAN, GEO MEAN	4.998, 4.431	0.009, 0.009	
	SD, SE	2.9163, 1.1906	NA, NA	
	MEDIAN	4.687	0.009	
	MIN, MAX	2.318, 10.545	0.009, 0.009	
	5%, 95%	2.318, 10.545	NA, NA	
	BLQ Values Included	0	0	
Week 3	N	5		NA
	MEAN, GEO MEAN	4.826, 2.324		
	SD, SE	6.5574, 2.9326		
	MEDIAN	2.547		
	MIN, MAX	0.609, 16.275		
	5%, 95%	0.609, 16.275		
	BLQ Values Included	0		
Overall[*]	N	6	2	1.0000
	MEAN, GEO MEAN	4.837, 3.842	0.032, 0.021	
	SD, SE	4.3685, 1.7834	0.0332, 0.0235	
	MEDIAN	3.397	0.032	
	MIN, MAX	2.192, 13.613	0.008, 0.055	
	5%, 95%	2.192, 13.613	0.008, 0.055	

[a] P-value from wilcoxon rank-sum test.
[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Total NNAL (pg/mL)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[*]
Week 1	N	49	66	0.0000
	MEAN, GEO MEAN	546.8, 341.2	9.5, 6.7	
	SD, SE	468.94, 66.99	22.61, 2.78	
	MEDIAN	475.2	6.0	
	MIN, MAX	6.0, 2045.7	6.0, 185.7	
	5%, 95%	16.5, 1604.2	6.0, 8.8	
	BLQ Values Included	1	59	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	552.8, 367.9	7.4, 6.6	
	SD, SE	487.35, 71.86	6.09, 0.75	
	MEDIAN	426.6	6.0	
	MIN, MAX	6.0, 2223.3	6.0, 47.3	
	5%, 95%	53.0, 1721.9	6.0, 11.9	
	BLQ Values Included	1	59	
Overall[*]	N	50	66	0.5348
	MEAN, GEO MEAN	542.5, 366.0	8.4, 6.8	
	SD, SE	427.04, 60.39	12.16, 1.50	
	MEDIAN	433.9	6.0	
	MIN, MAX	6.0, 1663.1	6.0, 95.9	
	5%, 95%	34.8, 1471.5	6.0, 11.9	

[*] P-value from wilcoxon rank-sum test.
[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Total NNAL (pmol/L)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	49	66	0.0000
	MEAN, GEO MEAN	2613.5, 1631.1	45.4, 31.8	
	SD, SE	2241.53, 320.22	108.05, 13.30	
	MEDIAN	2271.5	28.7	
	MIN, MAX	28.7, 9778.4	28.7, 887.6	
	5%, 95%	78.9, 7668.1	28.7, 42.1	
	BLQ Values Included	1	59	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	2642.5, 1758.6	35.2, 31.5	
	SD, SE	2329.54, 343.47	29.12, 3.58	
	MEDIAN	2038.9	28.7	
	MIN, MAX	28.7, 10627.4	28.7, 225.1	
	5%, 95%	253.3, 8230.7	28.7, 56.9	
	BLQ Values Included	1	59	
Overall[*]	N	50	66	0.5348
	MEAN, GEO MEAN	2592.9, 1749.4	40.3, 32.5	
	SD, SE	2041.22, 288.67	58.10, 7.15	
	MEDIAN	2074.0	28.7	
	MIN, MAX	28.7, 7949.4	28.7, 458.2	
	5%, 95%	166.1, 7033.6	28.7, 56.7	

[a] P-value from wilcoxon rank-sum test.
[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

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TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Free NNAL (pg/mL)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	150.0, 106.4	6.4, 6.2	
	SD, SE	107.23, 15.16	1.87, 0.23	
	MEDIAN	136.4	6.0	
	MIN, MAX	6.0, 484.5	6.0, 20.3	
	5%, 95%	11.4, 347.9	6.0, 8.5	
	BLQ Values Included	2	62	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	161.2, 123.5	7.3, 6.5	
	SD, SE	109.14, 16.09	6.26, 0.77	
	MEDIAN	144.2	6.0	
	MIN, MAX	6.0, 485.1	6.0, 42.4	
	5%, 95%	28.2, 405.1	6.0, 10.5	
	BLQ Values Included	1	62	
Overall[b]	N	50	66	0.1959
	MEAN, GEO MEAN	153.7, 118.3	6.8, 6.5	
	SD, SE	101.02, 14.29	3.24, 0.40	
	MEDIAN	138.9	6.0	
	MIN, MAX	6.0, 474.5	6.0, 24.2	
	5%, 95%	20.9, 351.6	6.0, 11.4	

[a] P-value from wilcoxon rank-sum test.
[b] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Free NNAL (pmol/L)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	717.2, 508.7	30.4, 29.8	
	SD, SE	512.56, 72.49	8.94, 1.10	
	MEDIAN	652.0	28.7	
	MIN, MAX	28.7, 2315.9	28.7, 97.0	
	5%, 95%	54.5, 1663.0	28.7, 40.6	
	BLQ Values Included	2	62	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	770.5, 590.3	34.9, 31.2	
	SD, SE	521.68, 76.92	29.94, 3.69	
	MEDIAN	689.0	28.7	
	MIN, MAX	28.7, 2318.8	28.7, 202.7	
	5%, 95%	134.8, 1936.4	28.7, 50.2	
	BLQ Values Included	1	62	
Overall[*]	N	50	66	0.1959
	MEAN, GEO MEAN	734.7, 565.3	32.7, 31.1	
	SD, SE	482.89, 68.29	15.45, 1.90	
	MEDIAN	663.9	28.7	
	MIN, MAX	28.7, 2268.1	28.7, 115.7	
	5%, 95%	99.9, 1680.7	28.7, 54.3	

[a] P-value from wilcoxon rank-sum test.
[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Total Nicotine (ng/mL)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	1612.6, 916.5	2.4, 1.4	
	SD, SE	1653.11, 233.78	4.47, 0.55	
	MEDIAN	1000.4	1.0	
	MIN, MAX	15.7, 8096.4	1.0, 23.2	
	5%, 95%	88.4, 5070.3	1.0, 8.8	
	BLQ Values Included	0	44	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	1794.7, 1094.9	3.3, 1.3	
	SD, SE	1855.53, 265.08	14.92, 1.84	
	MEDIAN	1304.2	1.0	
	MIN, MAX	11.9, 10733.1	1.0, 122.3	
	5%, 95%	151.2, 5120.6	1.0, 5.1	
	BLQ Values Included	0	51	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	1611.1, 1068.8	1.4, 1.2	
	SD, SE	1234.16, 181.97	1.86, 0.23	
	MEDIAN	1301.9	1.0	
	MIN, MAX	9.6, 4831.6	1.0, 15.2	
	5%, 95%	253.9, 3911.0	1.0, 2.1	
	BLQ Values Included	0	52	
Overall[*]	N	50	66	0.7780
	MEAN, GEO MEAN	1646.9, 1057.4	2.4, 1.4	
	SD, SE	1449.86, 205.04	6.08, 0.75	
	MEDIAN	1216.9	1.0	
	MIN, MAX	12.4, 7887.0	1.0, 48.6	
	5%, 95%	163.9, 3865.6	1.0, 4.5	

[a] P-value from wilcoxon rank-sum test.

[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Total Nicotine (nmol/L)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	9941.7, 5650.1	14.8, 8.8	
	SD, SE	10191.40, 1441.28	27.56, 3.39	
	MEDIAN	6167.2	6.2	
	MIN, MAX	96.8, 49914.3	6.2, 143.0	
	5%, 95%	545.0, 31258.4	6.2, 54.3	
	BLQ Values Included	0	44	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	11064.6, 6750.1	20.4, 8.2	
	SD, SE	11439.30, 1634.19	92.00, 11.32	
	MEDIAN	8040.4	6.2	
	MIN, MAX	73.4, 66169.6	6.2, 754.0	
	5%, 95%	932.1, 31568.5	6.2, 31.4	
	BLQ Values Included	0	51	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	9932.6, 6589.5	8.9, 7.4	
	SD, SE	7608.62, 1121.83	11.43, 1.41	
	MEDIAN	8025.9	6.2	
	MIN, MAX	59.2, 29786.8	6.2, 93.7	
	5%, 95%	1565.3, 24111.3	6.2, 12.9	
	BLQ Values Included	0	52	
Overall[b]	N	50	66	0.7780
	MEAN, GEO MEAN	10152.9, 6518.8	14.7, 8.6	
	SD, SE	8938.40, 1264.08	37.52, 4.62	
	MEDIAN	7502.2	6.2	
	MIN, MAX	76.5, 48623.6	6.2, 299.8	
	5%, 95%	1010.2, 23831.4	6.2, 28.0	

[a] P-value from wilcoxon rank-sum test.
[b] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Total Cotinine (ng/mL)

Timepoint	Statistic	Smokers	Non-Smokers	P-value*
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	3187.2, 2002.0	3.0, 1.8	
	SD, SE	2909.78, 411.50	5.07, 0.62	
	MEDIAN	2020.8	1.0	
	MIN, MAX	31.6, 12610.5	1.0, 35.5	
	5%, 95%	948.7, 10113.6	1.0, 10.7	
	BLQ Values Included	0	34	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	3644.7, 2368.7	3.4, 1.9	
	SD, SE	3403.71, 486.24	5.66, 0.70	
	MEDIAN	2690.9	1.1	
	MIN, MAX	28.5, 18373.7	1.0, 33.4	
	5%, 95%	503.9, 9222.8	1.0, 14.1	
	BLQ Values Included	0	33	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	3056.7, 2160.4	2.5, 1.6	
	SD, SE	2083.46, 307.19	4.40, 0.54	
	MEDIAN	2403.3	1.0	
	MIN, MAX	20.4, 7919.1	1.0, 33.8	
	5%, 95%	390.5, 6761.2	1.0, 6.2	
	BLQ Values Included	0	40	
Overall'	N	50	66	0.8968
	MEAN, GEO MEAN	3256.3, 2246.5	3.0, 1.9	
	SD, SE	2553.08, 361.06	4.28, 0.53	
	MEDIAN	2699.1	1.2	
	MIN, MAX	26.8, 12581.8	1.0, 27.1	
	5%, 95%	464.3, 7394.7	1.0, 11.3	

* P-value from wilcoxon rank-sum test.
' Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Total Cotinine (nmol/L)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	18087.2, 11361.6	17.1, 10.0	
	SD, SE	16513.00, 2335.29	28.76, 3.54	
	MEDIAN	11468.1	5.7	
	MIN, MAX	179.3, 71564.6	5.7, 201.5	
	5%, 95%	1978.9, 57394.7	5.7, 60.7	
	BLQ Values Included	0	34	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	20683.9, 13442.5	19.2, 10.6	
	SD, SE	19316.10, 2759.44	32.12, 3.95	
	MEDIAN	15270.9	6.3	
	MIN, MAX	161.7, 104270.7	5.7, 189.5	
	5%, 95%	2859.6, 52339.4	5.7, 80.0	
	BLQ Values Included	0	33	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	17347.0, 12260.6	14.1, 9.0	
	SD, SE	11823.60, 1743.30	24.95, 3.07	
	MEDIAN	13638.4	5.7	
	MIN, MAX	115.8, 44940.9	5.7, 191.8	
	5%, 95%	2216.1, 38369.8	5.7, 35.2	
	BLQ Values Included	0	40	
Overall*	N	50	66	0.8901
	MEAN, GEO MEAN	18479.7, 12749.4	16.8, 10.6	
	SD, SE	14488.80, 2049.02	24.26, 2.99	
	MEDIAN	15317.2	6.7	
	MIN, MAX	152.3, 71401.7	5.7, 153.6	
	5%, 95%	2634.7, 41965.1	5.7, 64.0	

[a] P-value from wilcoxon rank-sum test.
* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminating all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Free Nicotine (ng/mL)

Timepoint	Statistic	Smokers	Non-Smokers	P-value*
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	1141.8, 626.3	2.2, 1.3	
	SD, SE	1210.89, 171.25	4.22, 0.52	
	MEDIAN	649.3	1.0	
	MIN, MAX	12.8, 6329.6	1.0, 24.5	
	5%, 95%	45.8, 3752.0	1.0, 6.8	
	BLQ Values Included	0	52	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	1245.5, 756.4	2.9, 1.2	
	SD, SE	1251.38, 178.77	12.95, 1.59	
	MEDIAN	835.4	1.0	
	MIN, MAX	12.2, 6634.3	1.0, 106.2	
	5%, 95%	83.5, 3489.6	1.0, 4.3	
	BLQ Values Included	0	55	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	1166.7, 740.9	1.2, 1.1	
	SD, SE	933.84, 137.69	1.23, 0.15	
	MEDIAN	911.3	1.0	
	MIN, MAX	5.2, 3630.2	1.0, 10.5	
	5%, 95%	131.0, 3006.4	1.0, 1.7	
	BLQ Values Included	0	58	
Overall*	N	50	66	0.5895
	MEAN, GEO MEAN	1167.2, 745.0	2.1, 1.3	
	SD, SE	1017.73, 143.93	5.30, 0.65	
	MEDIAN	815.5	1.0	
	MIN, MAX	10.1, 5531.4	1.0, 42.7	
	5%, 95%	69.4, 2950.7	1.0, 3.3	

* P-value from wilcoxon rank-sum test.
* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminating all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Free Nicotine (nmol/L)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	7039.3, 3861.3	13.4, 8.1	
	SD, SE	7465.15, 1055.73	25.99, 3.20	
	MEDIAN	4002.7	6.2	
	MIN, MAX	78.9, 39022.0	6.2, 151.0	
	5%, 95%	282.4, 23131.1	6.2, 41.9	
	BLQ Values Included	0	52	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	7678.6, 4663.2	18.0, 7.7	
	SD, SE	7714.75, 1102.11	79.81, 9.82	
	MEDIAN	5150.2	6.2	
	MIN, MAX	75.2, 40900.5	6.2, 654.7	
	5%, 95%	514.8, 21513.4	6.2, 26.5	
	BLQ Values Included	0	55	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	7192.6, 4567.8	7.6, 6.8	
	SD, SE	5757.14, 848.84	7.55, 0.93	
	MEDIAN	5618.2	6.2	
	MIN, MAX	32.1, 22380.2	6.2, 64.7	
	5%, 95%	807.6, 18534.5	6.2, 10.5	
	BLQ Values Included	0	58	
Overall[b]	N	50	66	0.5875
	MEAN, GEO MEAN	7196.0, 4592.7	13.0, 8.0	
	SD, SE	6274.31, 887.32	32.67, 4.02	
	MEDIAN	5027.4	6.2	
	MIN, MAX	62.1, 34100.9	6.2, 263.2	
	5%, 95%	427.6, 18191.3	6.2, 20.3	

[a] P-value from wilcoxon rank-sum test.
[b] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

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TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Free Cotinine (ng/mL)

Timepoint	Statistic	Smokers	Non-Smokers	P-value*
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	1029.4, 773.3	1.7, 1.3	
	SD, SE	619.01, 87.54	2.08, 0.26	
	MEDIAN	919.4	1.0	
	MIN, MAX	18.2, 2626.8	1.0, 14.1	
	5%, 95%	226.0, 2263.4	1.0, 4.8	
	BLQ Values Included	0	50	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	1154.8, 883.0	1.9, 1.3	
	SD, SE	788.02, 112.57	2.89, 0.36	
	MEDIAN	975.4	1.0	
	MIN, MAX	15.5, 4778.6	1.0, 20.8	
	5%, 95%	221.4, 2343.3	1.0, 4.1	
	BLQ Values Included	0	48	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	1082.9, 837.2	1.5, 1.2	
	SD, SE	632.01, 93.19	2.40, 0.30	
	MEDIAN	998.4	1.0	
	MIN, MAX	6.1, 2858.1	1.0, 19.9	
	5%, 95%	257.2, 2346.7	1.0, 2.8	
	BLQ Values Included	0	51	
Overall*	N	50	66	0.6133
	MEAN, GEO MEAN	1076.8, 846.7	1.7, 1.3	
	SD, SE	616.60, 87.20	2.09, 0.26	
	MEDIAN	970.2	1.0	
	MIN, MAX	13.5, 3090.2	1.0, 13.9	
	5%, 95%	275.9, 1962.5	1.0, 4.1	

* P-value from wilcoxon rank-sum test.
* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Free Cotinine (nmol/L)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[*]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	5842.0, 4388.4	9.5, 7.3	
	SD, SE	3512.90, 496.80	11.80, 1.45	
	MEDIAN	5217.3	5.7	
	MIN, MAX	103.3, 14907.1	5.7, 80.0	
	5%, 95%	1282.6, 12844.8	5.7, 27.2	
	BLQ Values Included	0	50	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	6553.5, 5010.8	10.6, 7.6	
	SD, SE	4472.01, 638.86	16.37, 2.02	
	MEDIAN	5535.4	5.7	
	MIN, MAX	88.0, 27118.6	5.7, 118.0	
	5%, 95%	1256.4, 13298.2	5.7, 23.3	
	BLQ Values Included	0	48	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	6145.2, 4751.3	8.7, 6.8	
	SD, SE	3586.67, 528.83	13.63, 1.68	
	MEDIAN	5665.7	5.7	
	MIN, MAX	34.6, 16219.7	5.7, 112.9	
	5%, 95%	1459.6, 13317.5	5.7, 15.9	
	BLQ Values Included	0	51	
Overall[*]	N	50	66	0.6133
	MEAN, GEO MEAN	6110.8, 4805.3	9.6, 7.5	
	SD, SE	3499.18, 494.86	11.85, 1.46	
	MEDIAN	5505.8	5.7	
	MIN, MAX	76.8, 17537.1	5.7, 78.7	
	5%, 95%	1566.0, 11137.0	5.7, 23.5	

[*] P-value from wilcoxon rank-sum test.
[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Total 3-hydroxycotinine (ng/mL)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	4845.2, 3134.1	5.3, 2.4	
	SD, SE	4478.12, 633.30	12.32, 1.52	
	MEDIAN	3846.4	1.5	
	MIN, MAX	57.0, 26581.8	1.0, 96.1	
	5%, 95%	547.1, 12291.4	1.0, 18.0	
	BLQ Values Included	0	24	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	5978.4, 4009.9	7.6, 2.8	
	SD, SE	4841.56, 691.65	22.05, 2.71	
	MEDIAN	4802.7	2.0	
	MIN, MAX	53.8, 23327.3	1.0, 169.4	
	5%, 95%	807.0, 16284.7	1.0, 26.2	
	BLQ Values Included	0	16	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	5051.2, 3479.7	4.3, 2.2	
	SD, SE	4249.51, 626.56	9.12, 1.12	
	MEDIAN	4050.7	1.4	
	MIN, MAX	49.9, 23404.7	1.0, 69.1	
	5%, 95%	703.4, 12479.5	1.0, 15.6	
	BLQ Values Included	0	26	
Overall[b]	N	50	66	0.5447
	MEAN, GEO MEAN	5246.8, 3729.5	5.7, 2.7	
	SD, SE	3898.69, 551.36	11.46, 1.41	
	MEDIAN	4786.8	2.0	
	MIN, MAX	53.6, 18644.1	1.0, 66.3	
	5%, 95%	745.1, 11100.3	1.0, 18.6	

[a] P-value from wilcoxon rank-sum test.
[b] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Total 3-hydroxycotinine (nmol/L)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	25209.6, 16307.0	27.4, 12.5	
	SD, SE	23299.70, 3295.07	64.09, 7.89	
	MEDIAN	20012.6	7.8	
	MIN, MAX	296.6, 138305.1	5.2, 500.0	
	5%, 95%	2846.6, 63952.2	5.2, 93.7	
	BLQ Values Included	0	24	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	31105.5, 20863.4	39.7, 14.3	
	SD, SE	25190.70, 3598.67	114.71, 14.12	
	MEDIAN	24988.4	10.2	
	MIN, MAX	279.9, 121371.9	5.2, 881.4	
	5%, 95%	4198.8, 84729.3	5.2, 136.3	
	BLQ Values Included	0	16	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	26281.4, 18105.1	22.4, 11.3	
	SD, SE	22110.20, 3259.97	47.47, 5.84	
	MEDIAN	21075.5	7.3	
	MIN, MAX	259.6, 121774.7	5.2, 359.5	
	5%, 95%	3659.8, 64930.8	5.2, 81.2	
	BLQ Values Included	0	26	
Overall*	N	50	66	0.5465
	MEAN, GEO MEAN	27299.0, 19404.3	29.8, 14.1	
	SD, SE	20284.90, 2868.72	59.64, 7.34	
	MEDIAN	24905.7	10.2	
	MIN, MAX	278.7, 97005.2	5.2, 344.8	
	5%, 95%	3876.6, 57754.9	5.2, 96.8	

[a] P-value from wilcoxon rank-sum test.
* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

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TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Free 3-hydroxycotinine (ng/mL)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	3549.7, 2312.1	4.4, 2.2	
	SD, SE	3173.27, 448.77	9.90, 1.22	
	MEDIAN	3110.5	1.4	
	MIN, MAX	30.1, 17333.8	1.0, 77.2	
	5%, 95%	422.2, 9882.6	1.0, 16.4	
	BLQ Values Included	0	23	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	4688.4, 3050.3	6.1, 2.4	
	SD, SE	3986.05, 569.44	17.57, 2.16	
	MEDIAN	3435.7	1.7	
	MIN, MAX	26.9, 18865.3	1.0, 135.0	
	5%, 95%	692.4, 13308.1	1.0, 21.4	
	BLQ Values Included	0	18	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	3636.9, 2545.4	3.5, 1.9	
	SD, SE	2820.50, 415.86	7.27, 0.89	
	MEDIAN	3045.4	1.3	
	MIN, MAX	20.4, 15599.2	1.0, 56.0	
	5%, 95%	525.1, 8394.0	1.0, 10.9	
	BLQ Values Included	0	31	
Overall[b]	N	50	66	0.5728
	MEAN, GEO MEAN	3928.4, 2775.7	4.7, 2.4	
	SD, SE	2928.37, 414.13	9.23, 1.14	
	MEDIAN	3463.0	1.8	
	MIN, MAX	25.8, 13970.9	1.0, 52.6	
	5%, 95%	601.4, 8965.5	1.0, 14.3	

[a] P-value from wilcoxon rank-sum test.
[b] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

PM3003565310

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Free 3-hydroxycotinine (nmol/L)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	18469.2, 12030.0	22.8, 11.3	
	SD, SE	16510.50, 2334.94	51.53, 6.34	
	MEDIAN	16183.9	7.1	
	MIN, MAX	156.6, 90187.8	5.2, 401.7	
	5%, 95%	2196.7, 51419.2	5.2, 85.3	
	BLQ Values Included	0	23	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	24393.8, 15870.9	31.9, 12.4	
	SD, SE	20739.40, 2962.78	91.42, 11.25	
	MEDIAN	17875.9	8.8	
	MIN, MAX	140.0, 98156.2	5.2, 702.4	
	5%, 95%	3602.6, 69242.0	5.2, 111.3	
	BLQ Values Included	0	18	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	18922.7, 13243.4	18.3, 10.1	
	SD, SE	14675.10, 2163.72	37.80, 4.65	
	MEDIAN	15845.0	6.8	
	MIN, MAX	106.1, 81162.6	5.2, 291.4	
	5%, 95%	2732.1, 43674.0	5.2, 56.7	
	BLQ Values Included	0	31	
Overall[*]	N	50	66	0.5709
	MEAN, GEO MEAN	20439.3, 14441.8	24.4, 12.4	
	SD, SE	15236.30, 2154.74	48.03, 5.91	
	MEDIAN	18017.8	9.3	
	MIN, MAX	134.2, 72690.6	5.2, 273.7	
	5%, 95%	3129.3, 46647.3	5.2, 74.6	

[a] P-value from wilcoxon rank-sum test.

[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Total Nicotine (nmol/L)

Timepoint	Statistic	Smokers	Non-Smokers	P-value*
Week 1	N	6	3	0.0256
	MEAN, GEO MEAN	9798.9, 6501.7	12.7, 12.7	
	SD, SE	10412.60, 4250.94	0.00, 0.00	
	MEDIAN	4765.0	12.7	
	MIN, MAX	2468.7, 29112.4	12.7, 12.7	
	5%, 95%	2468.7, 29112.4	12.7, 12.7	
	BLQ Values Included	0	3	
Week 2	N	6	5	0.0067
	MEAN, GEO MEAN	8934.2, 6661.8	20.0, 16.7	
	SD, SE	7476.72, 3052.36	16.41, 7.34	
	MEDIAN	5769.3	12.7	
	MIN, MAX	2954.7, 21362.0	12.7, 49.4	
	5%, 95%	2954.7, 21362.0	12.7, 49.4	
	BLQ Values Included	0	4	
Week 3	N	5	7	0.0021
	MEAN, GEO MEAN	5251.1, 4536.8	12.7, 12.7	
	SD, SE	2902.43, 1298.01	0.00, 0.00	
	MEDIAN	5824.0	12.7	
	MIN, MAX	1973.7, 9133.4	12.7, 12.7	
	5%, 95%	1973.7, 9133.4	12.7, 12.7	
	BLQ Values Included	0	7	
Overall*	N	6	7	0.9401
	MEAN, GEO MEAN	8500.5, 6413.7	14.4, 14.0	
	SD, SE	7026.29, 2868.47	4.61, 1.74	
	MEDIAN	5421.7	12.7	
	MIN, MAX	2744.5, 19869.3	12.7, 24.9	
	5%, 95%	2744.5, 19869.3	12.7, 24.9	

* P-value from wilcoxon rank-sum test.
* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

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TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Total Nicotine (ng/mL)

Timepoint	Statistic	Smokers	Non-Smokers	P-value*
Week 1	N	6	3	0.0256
	MEAN, GEO MEAN	1589.4, 1054.6	2.1, 2.1	
	SD, SE	1688.92, 689.50	0.00, 0.00	
	MEDIAN	772.9	2.1	
	MIN, MAX	400.4, 4722.0	2.1, 2.1	
	5%, 95%	400.4, 4722.0	2.1, 2.1	
	BLQ Values Included	0	3	
Week 2	N	6	5	0.0067
	MEAN, GEO MEAN	1449.1, 1080.5	3.3, 2.7	
	SD, SE	1212.72, 495.09	2.64, 1.18	
	MEDIAN	935.8	2.1	
	MIN, MAX	479.3, 3464.9	2.1, 8.0	
	5%, 95%	479.3, 3464.9	2.1, 8.0	
	BLQ Values Included	0	4	
Week 3	N	5	7	0.0021
	MEAN, GEO MEAN	851.7, 735.9	2.1, 2.1	
	SD, SE	470.78, 210.54	0.00, 0.00	
	MEDIAN	944.7	2.1	
	MIN, MAX	320.1, 1481.4	2.1, 2.1	
	5%, 95%	320.1, 1481.4	2.1, 2.1	
	BLQ Values Included	0	7	
Overall*	N	6	7	0.9401
	MEAN, GEO MEAN	1378.8, 1040.3	2.4, 2.3	
	SD, SE	1139.66, 465.26	0.76, 0.29	
	MEDIAN	879.4	2.1	
	MIN, MAX	445.2, 3222.8	2.1, 4.1	
	5%, 95%	445.2, 3222.8	2.1, 4.1	

* P-value from wilcoxon rank-sum test.
* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 90 of ?)

Biomarker: Free Nicotine (nmol/L)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	3	0.0256
	MEAN, GEO MEAN	7197.1, 4444.0	3.4, 3.4	
	SD, SE	7757.37, 3166.93	0.00, 0.00	
	MEDIAN	3394.6	3.4	
	MIN, MAX	1356.6, 20738.8	3.4, 3.4	
	5%, 95%	1356.6, 20738.8	3.4, 3.4	
	BLQ Values Included	0	3	
Week 2	N	6	2	0.0651
	MEAN, GEO MEAN	6056.4, 4199.7	3.4, 3.4	
	SD, SE	5389.15, 2200.11	0.00, 0.00	
	MEDIAN	3774.8	3.4	
	MIN, MAX	1805.1, 14396.2	3.4, 3.4	
	5%, 95%	1805.1, 14396.2	3.4, 3.4	
	BLQ Values Included	0	2	
Week 3	N	5	4	0.0151
	MEAN, GEO MEAN	3400.4, 2972.7	3.4, 3.4	
	SD, SE	1754.96, 784.84	0.00, 0.00	
	MEDIAN	4113.9	3.4	
	MIN, MAX	1528.7, 5294.2	3.4, 3.4	
	5%, 95%	1528.7, 5294.2	3.4, 3.4	
	BLQ Values Included	0	4	
Overall[b]	N	6	6	0.7325
	MEAN, GEO MEAN	6005.1, 4360.7	3.4, 3.4	
	SD, SE	5159.10, 2106.19	0.00, 0.00	
	MEDIAN	3513.2	3.4	
	MIN, MAX	1564.8, 13476.4	3.4, 3.4	
	5%, 95%	1564.8, 13476.4	3.4, 3.4	

[a] P-value from wilcoxon rank-sum test.
[b] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 91 of ?)

Biomarker: Free Nicotine (ng/mL)

Timepoint	Statistic	Smokers	Non-Smokers	P-value*
Week 1	N	6	3	0.0256
	MEAN, GEO MEAN	1167.4, 720.8	0.6, 0.6	
	SD, SE	1258.24, 513.67	0.00, 0.00	
	MEDIAN	550.6	0.6	
	MIN, MAX	220.0, 3363.8	0.6, 0.6	
	5%, 95%	220.0, 3363.8	0.6, 0.6	
	BLQ Values Included	0	3	
Week 2	N	6	2	0.0651
	MEAN, GEO MEAN	982.4, 681.2	0.6, 0.6	
	SD, SE	874.12, 356.86	0.00, 0.00	
	MEDIAN	612.3	0.6	
	MIN, MAX	292.8, 2335.1	0.6, 0.6	
	5%, 95%	292.8, 2335.1	0.6, 0.6	
	BLQ Values Included	0	2	
Week 3	N	5	4	0.0151
	MEAN, GEO MEAN	551.6, 482.2	0.6, 0.6	
	SD, SE	284.63, 127.29	0.00, 0.00	
	MEDIAN	667.3	0.6	
	MIN, MAX	248.0, 858.7	0.6, 0.6	
	5%, 95%	248.0, 858.7	0.6, 0.6	
	BLQ Values Included	0	4	
Overall*	N	6	6	0.7325
	MEAN, GEO MEAN	974.0, 707.3	0.6, 0.6	
	SD, SE	836.80, 341.62	0.00, 0.00	
	MEDIAN	569.9	0.6	
	MIN, MAX	253.8, 2185.9	0.6, 0.6	
	5%, 95%	253.8, 2185.9	0.6, 0.6	

* P-value from Wilcoxon rank-sum test.
* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1,234' will be used as '1,234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

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TABLE 14.1.1-2

STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL

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Biomarker: Total Cotinine (nmol/L)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	6	0.0028
	MEAN, GEO MEAN	14356.4, 10535.4	11.0, 11.0	
	SD, SE	13129.60, 5360.14	0.00, 0.00	
	MEDIAN	11017.2	11.0	
	MIN, MAX	2865.9, 39806.6	11.0, 11.0	
	5%, 95%	2865.9, 39806.6	11.0, 11.0	
	BLQ Values Included	0	6	
Week 2	N	6	7	0.0015
	MEAN, GEO MEAN	14653.4, 11763.7	11.0, 11.0	
	SD, SE	10167.90, 4151.01	0.00, 0.00	
	MEDIAN	11093.9	11.0	
	MIN, MAX	3809.5, 30867.0	11.0, 11.0	
	5%, 95%	3809.5, 30867.0	11.0, 11.0	
	BLQ Values Included	0	7	
Week 3	N	5	7	0.0021
	MEAN, GEO MEAN	9971.7, 7688.8	11.0, 11.0	
	SD, SE	6819.34, 3049.70	0.00, 0.00	
	MEDIAN	8330.8	11.0	
	MIN, MAX	1766.5, 20274.4	11.0, 11.0	
	5%, 95%	1766.5, 20274.4	11.0, 11.0	
	BLQ Values Included	0	7	
Overall[b]	N	6	8	0.8875
	MEAN, GEO MEAN	13463.2, 10521.5	11.0, 11.0	
	SD, SE	9773.22, 3989.90	0.00, 0.00	
	MEDIAN	10588.5	11.0	
	MIN, MAX	2814.0, 30316.0	11.0, 11.0	
	5%, 95%	2814.0, 30316.0	11.0, 11.0	

[a] P-value from wilcoxon rank-sum test.

[b] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

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TABLE 14.1.1-2

STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Total Cotinine (ng/mL)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	6	0.0028
	MEAN, GEO MEAN	2529.6, 1856.3	1.9, 1.9	
	SD, SE	2313.43, 944.45	0.00, 0.00	
	MEDIAN	1941.2	1.9	
	MIN, MAX	505.0, 7013.9	1.9, 1.9	
	5%, 95%	505.0, 7013.9	1.9, 1.9	
	BLQ Values Included	0	6	
Week 2	N	6	7	0.0015
	MEAN, GEO MEAN	2581.9, 2072.8	1.9, 1.9	
	SD, SE	1791.59, 731.41	0.00, 0.00	
	MEDIAN	1954.8	1.9	
	MIN, MAX	671.2, 5438.8	1.9, 1.9	
	5%, 95%	671.2, 5438.8	1.9, 1.9	
	BLQ Values Included	0	7	
Week 3	N	5	7	0.0021
	MEAN, GEO MEAN	1757.0, 1354.8	1.9, 1.9	
	SD, SE	1201.54, 537.34	0.00, 0.00	
	MEDIAN	1467.9	1.9	
	MIN, MAX	311.3, 3572.3	1.9, 1.9	
	5%, 95%	311.3, 3572.3	1.9, 1.9	
	BLQ Values Included	0	7	
Overall[b]	N	6	8	0.8875
	MEAN, GEO MEAN	2372.2, 1853.9	1.9, 1.9	
	SD, SE	1722.05, 703.03	0.00, 0.00	
	MEDIAN	1865.7	1.9	
	MIN, MAX	495.8, 5341.7	1.9, 1.9	
	5%, 95%	495.8, 5341.7	1.9, 1.9	

[a] P-value from wilcoxon rank-sum test.

[b] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 94 of ?)

Biomarker: Free Cotinine (nmol/L)

Timepoint	Statistic	Smokers	Non-Smokers	P-value*
Week 1	N	6	7	0.0021
	MEAN, GEO MEAN	5138.7, 4383.0	5.6, 4.1	
	SD, SE	3563.54, 1454.81	6.73, 2.54	
	MEDIAN	4048.8	3.1	
	MIN, MAX	2279.8, 12007.9	3.1, 20.9	
	5%, 95%	2279.8, 12007.9	3.1, 20.9	
	BLQ Values Included	0	6	
Week 2	N	6	8	0.0008
	MEAN, GEO MEAN	5341.9, 4848.5	3.1, 3.1	
	SD, SE	2622.04, 1070.44	0.00, 0.00	
	MEDIAN	4597.2	3.1	
	MIN, MAX	2527.5, 9702.2	3.1, 3.1	
	5%, 95%	2527.5, 9702.2	3.1, 3.1	
	BLQ Values Included	0	8	
Week 3	N	5	7	0.0021
	MEAN, GEO MEAN	4021.5, 3476.2	3.1, 3.1	
	SD, SE	2413.87, 1079.52	0.00, 0.00	
	MEDIAN	3087.7	3.1	
	MIN, MAX	1520.8, 7780.3	3.1, 3.1	
	5%, 95%	1520.8, 7780.3	3.1, 3.1	
	BLQ Values Included	0	7	
Overall"	N	6	8	0.6823
	MEAN, GEO MEAN	4959.3, 4381.0	3.8, 3.5	
	SD, SE	2782.63, 1136.00	2.09, 0.74	
	MEDIAN	4206.5	3.1	
	MIN, MAX	2109.4, 9830.1	3.1, 9.0	
	5%, 95%	2109.4, 9830.1	3.1, 9.0	

* P-value from wilcoxon rank-sum test.
" Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

Source: https://www.industrydocuments.ucsf.edu/docs/zhnj0001

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 95 of ?)

Biomarker: Free Cotinine (ng/mL)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	7	0.0021
	MEAN, GEO MEAN	905.4, 772.3	1.0, 0.7	
	SD, SE	627.91, 256.34	1.21, 0.46	
	MEDIAN	713.4	0.5	
	MIN, MAX	401.7, 2115.8	0.5, 3.7	
	5%, 95%	401.7, 2115.8	0.5, 3.7	
	BLQ Values Included	0	6	
Week 2	N	6	8	0.0008
	MEAN, GEO MEAN	941.2, 854.3	0.5, 0.5	
	SD, SE	462.01, 188.61	0.00, 0.00	
	MEDIAN	810.0	0.5	
	MIN, MAX	445.3, 1709.5	0.5, 0.5	
	5%, 95%	445.3, 1709.5	0.5, 0.5	
	BLQ Values Included	0	8	
Week 3	N	5	7	0.0021
	MEAN, GEO MEAN	708.6, 612.5	0.5, 0.5	
	SD, SE	425.31, 190.20	0.00, 0.00	
	MEDIAN	544.1	0.5	
	MIN, MAX	268.0, 1370.9	0.5, 0.5	
	5%, 95%	268.0, 1370.9	0.5, 0.5	
	BLQ Values Included	0	7	
Overall[b]	N	6	8	0.6823
	MEAN, GEO MEAN	873.9, 771.9	0.6, 0.6	
	SD, SE	490.31, 200.17	0.39, 0.14	
	MEDIAN	741.2	0.5	
	MIN, MAX	371.7, 1732.1	0.5, 1.6	
	5%, 95%	371.7, 1732.1	0.5, 1.6	

[a] P-value from wilcoxon rank-sum test.

[b] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 96 of ?)

Biomarker: Total 3-hydroxycotinine (nmol/L)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	5	0.0075
	MEAN, GEO MEAN	18760.8, 17494.6	24.5, 20.3	
	SD, SE	6817.90, 2783.39	16.64, 7.44	
	MEDIAN	20361.6	12.4	
	MIN, MAX	8417.0, 25578.5	12.4, 44.5	
	5%, 95%	8417.0, 25578.5	12.4, 44.5	
	BLQ Values Included	0	3	
Week 2	N	6	5	0.0067
	MEAN, GEO MEAN	19977.0, 17594.2	18.3, 15.8	
	SD, SE	10727.60, 4379.54	13.19, 5.90	
	MEDIAN	18523.4	12.4	
	MIN, MAX	7560.8, 37689.3	12.4, 41.9	
	5%, 95%	7560.8, 37689.3	12.4, 41.9	
	BLQ Values Included	0	4	
Week 3	N	5	5	0.0075
	MEAN, GEO MEAN	12925.5, 10547.5	12.4, 12.4	
	SD, SE	9343.85, 4178.70	0.00, 0.00	
	MEDIAN	9437.8	12.4	
	MIN, MAX	4279.9, 27954.4	12.4, 12.4	
	5%, 95%	4279.9, 27954.4	12.4, 12.4	
	BLQ Values Included	0	5	
Overall[b]	N	6	6	1.0000
	MEAN, GEO MEAN	18260.5, 16211.6	20.6, 17.8	
	SD, SE	9101.47, 3715.66	12.90, 5.27	
	MEDIAN	17343.5	12.4	
	MIN, MAX	6752.6, 31633.9	12.4, 40.9	
	5%, 95%	6752.6, 31633.9	12.4, 40.9	

[a] P-value from wilcoxon rank-sum test.
[b] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 97 of ?)

Biomarker: Total 3-hydroxycotinine (ng/mL)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	5	0.0075
	MEAN, GEO MEAN	3605.8, 3362.4	4.7, 3.9	
	SD, SE	1310.43, 534.98	3.21, 1.44	
	MEDIAN	3913.5	2.4	
	MIN, MAX	1617.7, 4916.2	2.4, 8.6	
	5%, 95%	1617.7, 4916.2	2.4, 8.6	
	BLQ Values Included	0	3	
Week 2	N	6	5	0.0067
	MEAN, GEO MEAN	3839.6, 3381.6	3.5, 3.1	
	SD, SE	2061.85, 841.75	2.55, 1.14	
	MEDIAN	3560.2	2.4	
	MIN, MAX	1453.2, 7243.9	2.4, 8.1	
	5%, 95%	1453.2, 7243.9	2.4, 8.1	
	BLQ Values Included	0	4	
Week 3	N	5	5	0.0075
	MEAN, GEO MEAN	2484.3, 2027.2	2.4, 2.4	
	SD, SE	1795.88, 803.14	0.00, 0.00	
	MEDIAN	1813.9	2.4	
	MIN, MAX	822.6, 5372.8	2.4, 2.4	
	5%, 95%	822.6, 5372.8	2.4, 2.4	
	BLQ Values Included	0	5	
Overall[*]	N	6	6	1.0000
	MEAN, GEO MEAN	3509.7, 3115.9	4.0, 3.4	
	SD, SE	1749.33, 714.16	2.50, 1.02	
	MEDIAN	3333.4	2.4	
	MIN, MAX	1297.8, 6080.1	2.4, 7.9	
	5%, 95%	1297.8, 6080.1	2.4, 7.9	

[a] P-value from wilcoxon rank-sum test.
[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

PM3003565321

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Free 3-hydroxycotinine (nmol/L)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	6	8	0.0024
	MEAN, GEO MEAN	15531.5, 14510.3	12.9, 10.0	
	SD, SE	5703.03, 2328.25	9.94, 3.51	
	MEDIAN	16253.2	8.3	
	MIN, MAX	7282.3, 21708.6	4.9, 31.9	
	5%, 95%	7282.3, 21708.6	4.9, 31.9	
	BLQ Values Included	0	0	
Week 2	N	6	7	0.0034
	MEAN, GEO MEAN	16215.6, 13989.4	12.7, 10.9	
	SD, SE	8993.85, 3671.72	7.86, 2.97	
	MEDIAN	14730.5	10.3	
	MIN, MAX	5006.9, 30541.0	6.0, 27.6	
	5%, 95%	5006.9, 30541.0	6.0, 27.6	
	BLQ Values Included	0	0	
Week 3	N	5	5	0.0122
	MEAN, GEO MEAN	10483.2, 8939.9	11.3, 10.7	
	SD, SE	6664.13, 2980.29	4.09, 1.83	
	MEDIAN	8860.2	12.4	
	MIN, MAX	4009.9, 21054.0	7.0, 16.9	
	5%, 95%	4009.9, 21054.0	7.0, 16.9	
	BLQ Values Included	0	0	
Overall*	N	6	8	1.0000
	MEAN, GEO MEAN	14945.7, 13284.5	11.3, 10.0	
	SD, SE	7410.05, 3025.14	6.05, 2.14	
	MEDIAN	14143.7	10.6	
	MIN, MAX	5433.0, 26124.8	5.5, 22.2	
	5%, 95%	5433.0, 26124.8	5.5, 22.2	

[a] P-value from wilcoxon rank-sum test.
* Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminating all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 99 of ?)

Biomarker: Free 3-hydroxycotinine (ng/mL)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[*]
Week 1	N	6	8	0.0024
	MEAN, GEO MEAN	2985.2, 2788.9	2.5, 1.9	
	SD, SE	1096.11, 447.48	1.92, 0.68	
	MEDIAN	3123.9	1.6	
	MIN, MAX	1399.7, 4172.4	0.9, 6.1	
	5%, 95%	1399.7, 4172.4	0.9, 6.1	
	BLQ Values Included	0	0	
Week 2	N	6	7	0.0034
	MEAN, GEO MEAN	3116.6, 2688.7	2.4, 2.1	
	SD, SE	1728.63, 705.71	1.50, 0.57	
	MEDIAN	2831.2	2.0	
	MIN, MAX	962.3, 5870.0	1.2, 5.3	
	5%, 95%	962.3, 5870.0	1.2, 5.3	
	BLQ Values Included	0	0	
Week 3	N	5	5	0.0119
	MEAN, GEO MEAN	2014.9, 1718.2	2.1, 2.0	
	SD, SE	1280.85, 572.81	0.79, 0.35	
	MEDIAN	1702.9	2.4	
	MIN, MAX	770.7, 4046.6	1.3, 3.2	
	5%, 95%	770.7, 4046.6	1.3, 3.2	
	BLQ Values Included	0	0	
Overall[*]	N	6	8	1.0000
	MEAN, GEO MEAN	2872.6, 2553.3	2.2, 1.9	
	SD, SE	1424.22, 581.44	1.15, 0.40	
	MEDIAN	2718.4	2.0	
	MIN, MAX	1044.2, 5021.2	1.1, 4.2	
	5%, 95%	1044.2, 5021.2	1.1, 4.2	

[*] P-value from wilcoxon rank-sum test.
[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

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TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Nicotine Glucuronide (nmol/L)

Timepoint	Statistic	Smokers	Non-Smokers	P-value*
Week 1	N	49	63	0.0000
	MEAN, GEO MEAN	2978.210, 1359.260	1.781, NA	
	SD, SE	3802.1400, 543.1640	4.2499, 0.5354	
	MEDIAN	1405.000	0.000	
	MIN, MAX	17.879, 17624.502	0.000, 22.811	
	5%, 95%	71.514, 11547.000	0.000, 9.248	
	BLQ Values Included	0	52	
Week 2	N	48	64	0.0000
	MEAN, GEO MEAN	3456.560, 1750.150	2.553, NA	
	SD, SE	4353.5900, 628.3870	12.5617, 1.5702	
	MEDIAN	1479.290	0.000	
	MIN, MAX	68.432, 25269.102	0.000, 99.257	
	5%, 95%	121.451, 10055.100	0.000, 9.864	
	BLQ Values Included	0	55	
Week 3	N	45	66	0.0000
	MEAN, GEO MEAN	2802.100, 1441.500	1.308, NA	
	SD, SE	2424.2900, 361.3920	4.0615, 0.4999	
	MEDIAN	2594.230	0.000	
	MIN, MAX	12.330, 8494.754	0.000, 28.976	
	5%, 95%	27.126, 8015.730	0.000, 6.165	
	BLQ Values Included	0	58	
Overall⁺	N	50	66	0.8154
	MEAN, GEO MEAN	2971.560, 1586.550	1.885, NA	
	SD, SE	3164.2200, 447.4880	5.3651, 0.6604	
	MEDIAN	1558.720	0.000	
	MIN, MAX	22.503, 14522.700	0.000, 36.579	
	5%, 95%	98.640, 9135.910	0.000, 7.707	

* P-value from wilcoxon rank-sum test.
⁺ Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1,234' will be used as '1,234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

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TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Nicotine Gluruonide (nmol/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	49	63	0.0000
	MEAN, GEO MEAN	4098.500, 2293.980	2.994, NA	
	SD, SE	4073.6500, 581.9500	6.9750, 0.8788	
	MEDIAN	2740.960	0.000	
	MIN, MAX	25.209, 22030.628	0.000, 29.594	
	5%, 95%	162.756, 10788.300	0.000, 22.042	
	BLQ Values Included	0	52	
Week 2	N	48	64	0.0000
	MEAN, GEO MEAN	4409.210, 2738.290	6.248, NA	
	SD, SE	4279.9900, 617.7630	35.4753, 4.4344	
	MEDIAN	3529.190	0.000	
	MIN, MAX	177.318, 22489.501	0.000, 282.882	
	5%, 95%	294.258, 11807.500	0.000, 14.886	
	BLQ Values Included	0	55	
Week 3	N	45	66	0.0000
	MEAN, GEO MEAN	4437.910, 2471.850	2.290, NA	
	SD, SE	3364.7100, 501.5820	6.1951, 0.7626	
	MEDIAN	4005.330	0.000	
	MIN, MAX	17.262, 12487.288	0.000, 37.669	
	5%, 95%	52.296, 10485.800	0.000, 14.303	
	BLQ Values Included	0	58	
Overall[b]	N	50	66	0.9128
	MEAN, GEO MEAN	4172.890, 2585.250	3.838, NA	
	SD, SE	3505.6200, 495.7700	13.4760, 1.6588	
	MEDIAN	3084.910	0.000	
	MIN, MAX	26.819, 15441.800	0.000, 104.041	
	5%, 95%	199.791, 11219.200	0.000, 18.618	

[a] P-value from wilcoxon rank-sum test.
[b] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

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TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
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Biomarker: Nicotine Glumonide (nmol/mg Creatin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	49	63	0.0000
	MEAN, GEO MEAN	3.019, 1.886	0.002, NA	
	SD, SE	2.3754, 0.3393	0.0047, 0.0006	
	MEDIAN	2.774	0.000	
	MIN, MAX	0.024, 12.113	0.000, 0.023	
	5%, 95%	0.135, 7.058	0.000, 0.012	
	BLQ Values Included	0	52	
Week 2	N	48	64	0.0000
	MEAN, GEO MEAN	3.238, 2.097	0.005, NA	
	SD, SE	3.1048, 0.4481	0.0338, 0.0042	
	MEDIAN	2.471	0.000	
	MIN, MAX	0.153, 17.646	0.000, 0.270	
	5%, 95%	0.222, 6.805	0.000, 0.011	
	BLQ Values Included	0	55	
Week 3	N	45	66	0.0000
	MEAN, GEO MEAN	3.372, 1.946	0.001, NA	
	SD, SE	2.5882, 0.3858	0.0034, 0.0004	
	MEDIAN	2.840	0.000	
	MIN, MAX	0.020, 10.645	0.000, 0.021	
	5%, 95%	0.056, 8.031	0.000, 0.008	
	BLQ Values Included	0	58	
Overall[b]	N	50	66	0.7949
	MEAN, GEO MEAN	3.169, 2.071	0.003, NA	
	SD, SE	2.4360, 0.3445	0.0121, 0.0015	
	MEDIAN	2.648	0.000	
	MIN, MAX	0.025, 10.339	0.000, 0.097	
	5%, 95%	0.190, 8.383	0.000, 0.009	

[a] P-value from wilcoxon rank-sum test.
[b] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

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TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 103 of ?)

Biomarker: Cotinine Gluruonide (nmol/L)

Timepoint	Statistic	Smokers	Non Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	12245.200, 6153.630	7.575, NA	
	SD, SE	13823.3000, 1954.9100	18.0877, 2.2264	
	MEDIAN	6149.150	0.000	
	MIN, MAX	71.505, 58435.475	0.000, 121.445	
	5%, 95%	193.518, 44549.900	0.000, 38.023	
	BLQ Values Included	0	50	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	14130.400, 7525.660	8.685, NA	
	SD, SE	15556.5000, 2222.3600	18.1773, 2.2375	
	MEDIAN	7567.610	0.568	
	MIN, MAX	73.775, 77152.193	0.000, 92.503	
	5%, 95%	310.423, 42264.000	0.000, 57.318	
	BLQ Values Included	0	48	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	11201.800, 6697.690	5.469, NA	
	SD, SE	9178.1500, 1353.2400	12.1389, 1.4942	
	MEDIAN	8706.590	0.000	
	MIN, MAX	81.153, 37456.135	0.000, 78.883	
	5%, 95%	756.478, 27258.700	0.000, 22.700	
	BLQ Values Included	0	51	
Overall[b]	N	50	66	0.8924
	MEAN, GEO MEAN	12368.900, 7152.220	7.243, NA	
	SD, SE	11679.4000, 1651.7200	13.5374, 1.6663	
	MEDIAN	8028.620	0.852	
	MIN, MAX	75.478, 53864.600	0.000, 74.910	
	5%, 95%	430.165, 31626.000	0.000, 33.293	

[a] P-value from wilcoxon rank-sum test.

[b] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 104 of ?)

Biomarker: Cotinine Glucuonide (nmol/24hr)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	17584.200, 10515.700	12.042, NA	
	SD, SE	15317.2000, 2166.1700	25.1101, 3.0908	
	MEDIAN	13520.000	0.000	
	MIN, MAX	100.822, 68661.683	0.000, 106.872	
	5%, 95%	561.202, 52173.800	0.000, 75.621	
	BLQ Values Included	0	50	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	18182.400, 11642.600	15.112, NA	
	SD, SE	14562.1000, 2080.3000	32.1430, 3.9565	
	MEDIAN	14973.800	0.698	
	MIN, MAX	66.398, 68665.452	0.000, 203.789	
	5%, 95%	523.023, 54491.400	0.000, 73.080	
	BLQ Values Included	0	48	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	17809.300, 11430.900	9.802, NA	
	SD, SE	12417.3000, 1830.8400	21.1127, 2.5988	
	MEDIAN	16848.200	0.000	
	MIN, MAX	85.048, 48692.976	0.000, 105.328	
	5%, 95%	1043.940, 37176.400	0.000, 50.216	
	BLQ Values Included	0	51	
Overall[b]	N	50	66	0.9708
	MEAN, GEO MEAN	17588.200, 11451.600	12.319, NA	
	SD, SE	13197.5000, 1866.4100	22.5981, 2.7816	
	MEDIAN	15932.300	1.638	
	MIN, MAX	84.089, 57165.200	0.000, 104.966	
	5%, 95%	827.405, 46579.300	0.000, 50.161	

[a] P-value from wilcoxon rank-sum test.
[b] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.

TABLE 14.1.1-2
STATISTICAL COMPARISON BETWEEN SMOKERS AND NON-SMOKERS
OF 24-HOUR URINE BIOMARKERS OF EXPOSURE BY TIMEPOINT AND OVERALL
(Page 105 of ?)

Biomarker: Cotinine Gluruonide (nmol/mg Creatin)

Timepoint	Statistic	Smokers	Non-Smokers	P-value[a]
Week 1	N	50	66	0.0000
	MEAN, GEO MEAN	13.335, 8.713	0.008, NA	
	SD, SE	9.4621, 1.3381	0.0173, 0.0021	
	MEDIAN	12.095	0.000	
	MIN, MAX	0.095, 37.108	0.000, 0.075	
	5%, 95%	0.507, 33.762	0.000, 0.055	
	BLQ Values Included	0	50	
Week 2	N	49	66	0.0000
	MEAN, GEO MEAN	13.437, 8.986	0.011, NA	
	SD, SE	10.4751, 1.4964	0.0267, 0.0033	
	MEDIAN	11.760	0.001	
	MIN, MAX	0.074, 53.877	0.000, 0.195	
	5%, 95%	0.522, 34.914	0.000, 0.043	
	BLQ Values Included	0	48	
Week 3	N	46	66	0.0000
	MEAN, GEO MEAN	13.808, 9.166	0.006, NA	
	SD, SE	10.2141, 1.5060	0.0143, 0.0018	
	MEDIAN	14.106	0.000	
	MIN, MAX	0.075, 49.743	0.000, 0.067	
	5%, 95%	1.668, 34.847	0.000, 0.035	
	BLQ Values Included	0	51	
Overall[*]	N	50	66	0.6097
	MEAN, GEO MEAN	13.481, 9.180	0.008, NA	
	SD, SE	9.5809, 1.3549	0.0166, 0.0020	
	MEDIAN	12.611	0.001	
	MIN, MAX	0.081, 38.906	0.000, 0.091	
	5%, 95%	0.809, 38.061	0.000, 0.047	

[a] P-value from wilcoxon rank-sum test.

[*] Overall values calculated by first averaging the results across visits within a subject, then calculating the descriptive statistics across subjects. Count of BLQ values does not apply as the BLQ values were converted to the upper limit of the BLQ (e.g., '<1.234' will be used as '1.234') for the averages across visits within a subject thus eliminatig all BLQ values for the overall statistics.